

Network Systems
Science & Advanced
Computing
Biocomplexity Institute
& Initiative
University of Virginia

Estimation of COVID-19 Impact in Virginia

November 30th, 2022

(data current to November 26th – November 29th)

Biocomplexity Institute Technical report: TR BI-2022-1781



BIOCOMPLEXITY INSTITUTE



biocomplexity.virginia.edu

About Us

- Biocomplexity Institute at the University of Virginia
 - Using big data and simulations to understand massively interactive systems and solve societal problems
- Over 20 years of crafting and analyzing infectious disease models
 - Pandemic response for Influenza, Ebola, Zika, and others



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Overview

- **Goal:** Understand impact of COVID-19 mitigations in Virginia
- **Approach:**
 - Calibrate explanatory mechanistic model to observed cases
 - Project based on scenarios for next 4 months
 - Consider a range of possible mitigation effects in "what-if" scenarios
- **Outcomes:**
 - Ill, Confirmed, Hospitalized, ICU, Ventilated, Death
 - Geographic spread over time, case counts, healthcare burdens

Key Takeaways

Projecting future cases precisely is impossible and unnecessary.

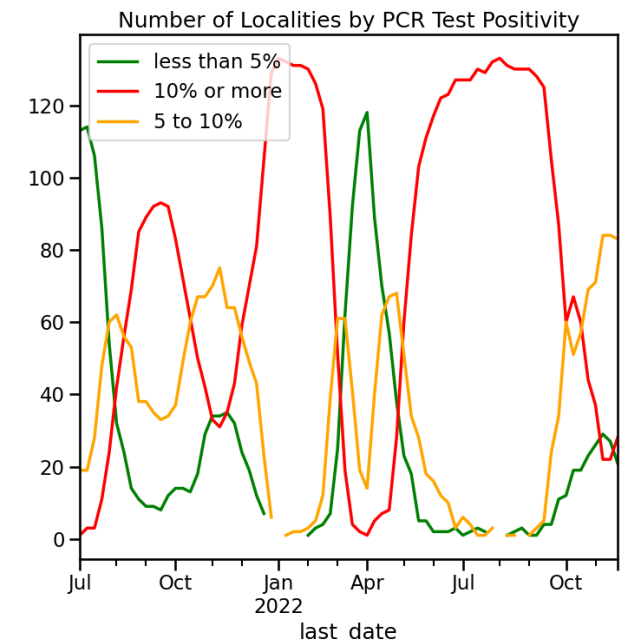
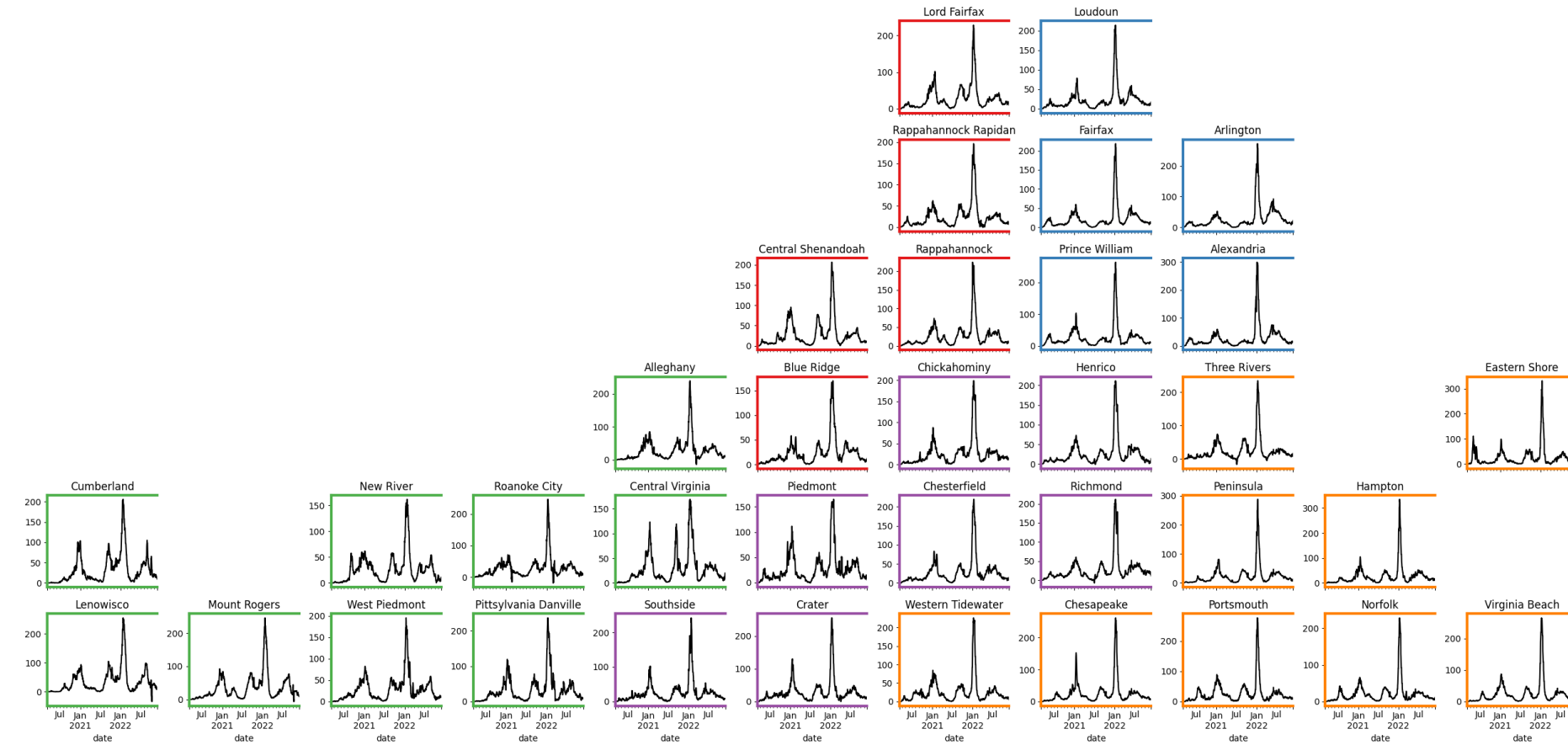
Even without perfect projections, we can confidently draw conclusions:

- **Case rates remain relatively flat though hospitalizations have experienced burst of growth in past week**
- VA weekly case rate is steady at 73/100K from 72/100K
 - US weekly case rate is flat remaining at 76 per 100K from 73 per 100K, though hospitalizations continue a steady rise
- VA hospital occupancy is quickly rising (rolling 7 day mean of 488 from 464 a week ago)
 - Influenza weekly hospital admissions remain high (~300 a week) and are approaching parity with COVID admissions (~40% of the combined)
- Projections anticipate increases in cases and hospitalizations in coming weeks
 - Combined hospitalizations due to Influenza and COVID-19 though are expected to level off
- Model updates:
 - Variant X candidates have now surpassed 50% (BQ.1.1, BN.1, BF.7 and others and XBB among others), 50% remains at Nov 12th
 - Modified Booster Scenarios: Current pace (included in all scenarios) with comparisons between keeping the pace (which is slowing) and a more Pessimistic scenario where vaccination halts at current levels

The situation continues to change. Models continue to be updated regularly.

Situation Assessment

Case Rates (per 100k) and Test Positivity



County level RT-PCR test positivity

Green: <5.0% (or <20 tests in past 14 days)
Orange: 5.0%-10.0% (or <500 tests and <2000 tests/100k and >10% positivity over 14 days)
Red: >10.0% (and not "Green" or "Yellow")

District Trajectories

Goal: Define epochs of a Health District's COVID-19 incidence to characterize the current trajectory

Method: Find recent peak and use hockey stick fit to find inflection point afterwards, then use this period's slope to define the trajectory

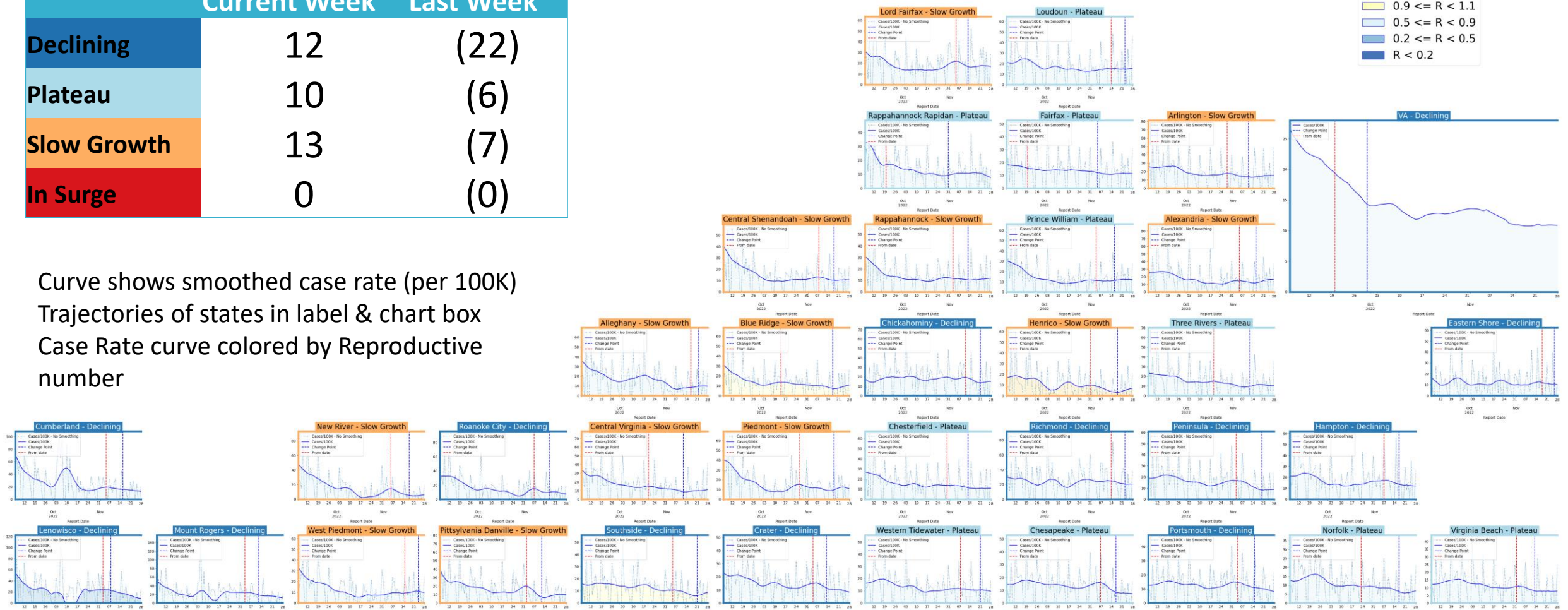
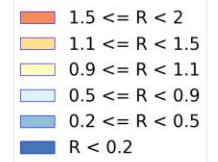


Trajectory	Description	Weekly Case Rate Slope (per 100k)	Weekly Hosp Rate Slope (per 100k)
Declining	Sustained decreases following a recent peak	$\text{slope} < -0.88/\text{day}$	$\text{slope} < -0.07/\text{day}$
Plateau	Steady level with minimal trend up or down	$-0.88/\text{day} < \text{slope} < 0.42/\text{day}$	$-0.07/\text{day} < \text{slope} < 0.07/\text{day}$
Slow Growth	Sustained growth not rapid enough to be considered a Surge	$0.42/\text{day} < \text{slope} < 2.45/\text{day}$	$0.07/\text{day} < \text{slope} < 0.21/\text{day}$
In Surge	Currently experiencing sustained rapid and significant growth	$2.45/\text{day} < \text{slope}$	$0.21/\text{day} < \text{slope}$

District Case Trajectories – last 10 weeks

Status	Number of Districts	
	Current Week	Last Week
Declining	12	(22)
Plateau	10	(6)
Slow Growth	13	(7)
In Surge	0	(0)

Curve shows smoothed case rate (per 100K)
 Trajectories of states in label & chart box
 Case Rate curve colored by Reproductive number



District Hospital Trajectories – last 10 weeks

Status	Number of Districts	
	Current Week	Last Week
Declining	18	(23)
In Surge	1	(3)
Plateau	12	(10)
Slow Growth	4	(0)

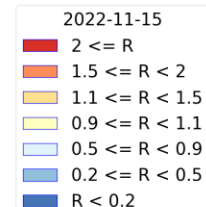
Holiday reporting delay

Hospitalization by county is delayed, these data are current as of

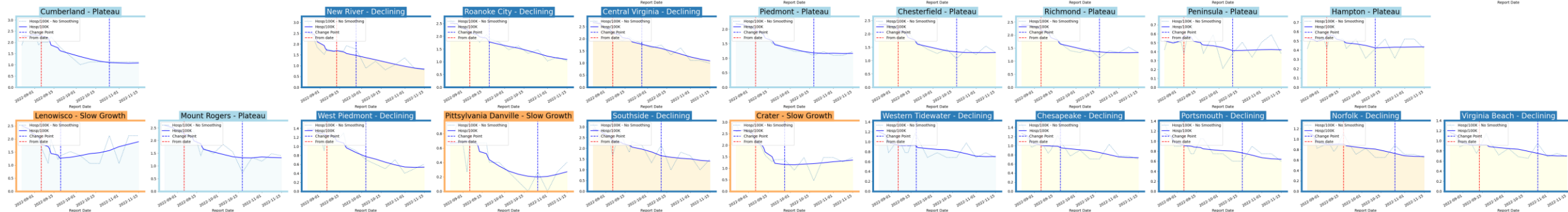
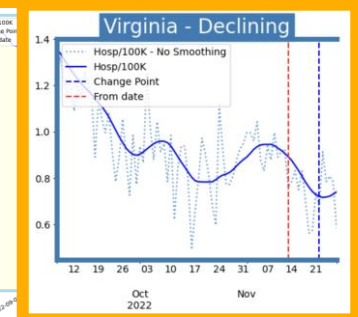
November 15th

Curve shows smoothed hospitalization rate (per 100K) by district

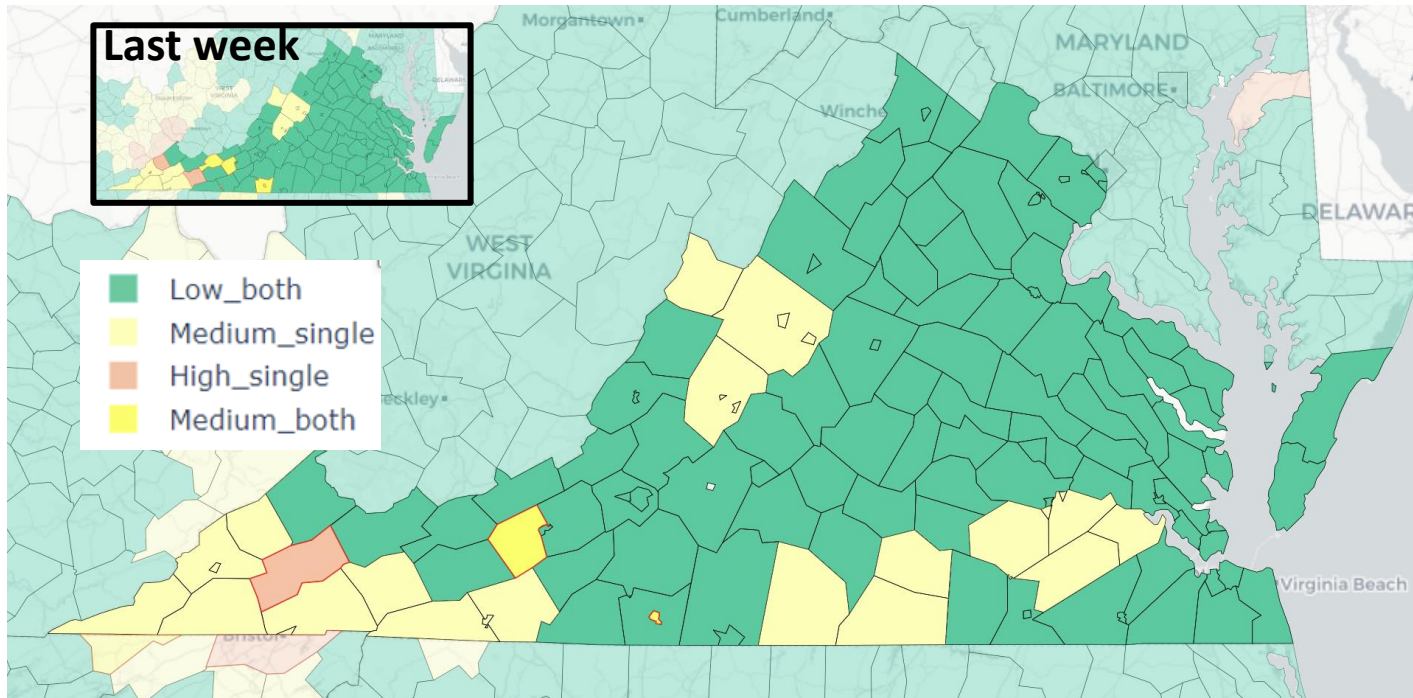
Hosp rate curve colored by R_e number



State level without delay



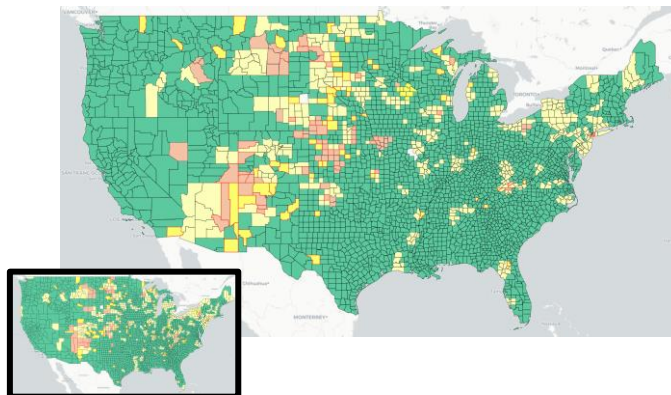
CDC's COVID-19 Community Levels



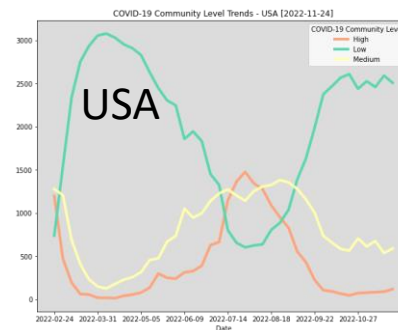
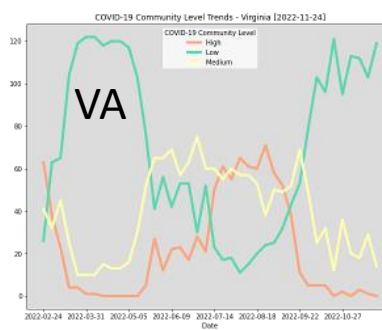
Red outline indicates county had 200 or more cases per 100k in last week

Pale color indicates either beds or occupancy set the level for this county

Dark color indicates both beds and occupancy set the level for this county



1-Dec-22



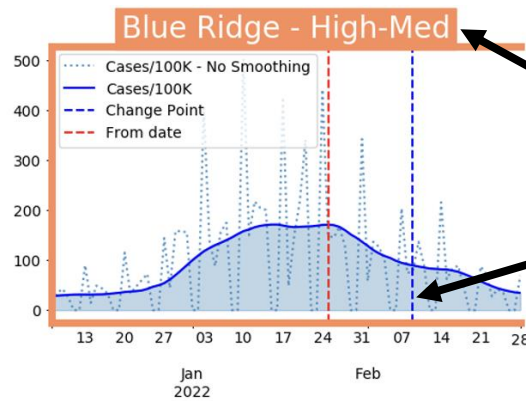
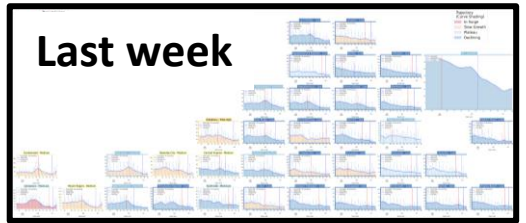
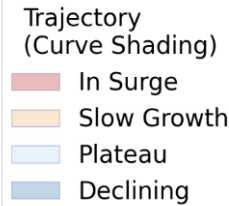
COVID-19 Community Levels – Use the Highest Level that Applies to Your Community				
New COVID-19 Cases Per 100,000 people in the past 7 days	Indicators	Low	Medium	High
Fewer than 200	New COVID-19 admissions per 100,000 population (7-day total)	<10.0	10.0-19.9	≥20.0
	Percent of staffed inpatient beds occupied by COVID-19 patients (7-day average)	<10.0%	10.0-14.9%	≥15.0%
200 or more	New COVID-19 admissions per 100,000 population (7-day total)	NA	<10.0	≥10.0
	Percent of staffed inpatient beds occupied by COVID-19 patients (7-day average)	NA	<10.0%	≥10.0%

The COVID-19 community level is determined by the higher of the new admissions and inpatient beds metrics, based on the current level of new cases per 100,000 population in the past 7 days

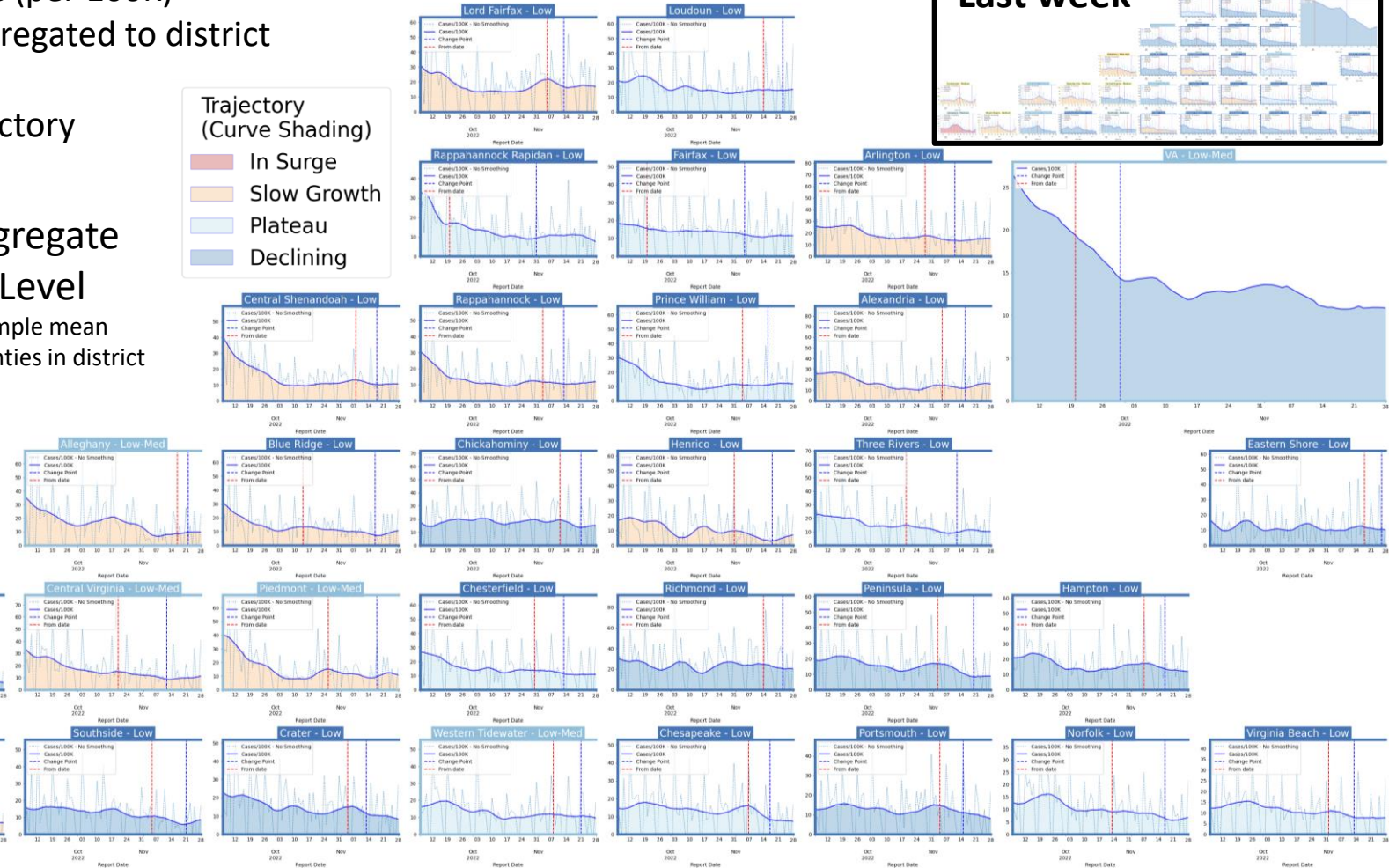
District Trajectories with Community Levels



Curve shows smoothed case rate (per 100K)
CDC's new [Community Level](#) aggregated to district level in label & chart box color
Case Rate curve colored by Trajectory



District's Aggregate
Community Level
Aggregate level a simple mean
of all levels for counties in district
Case rate
Trajectory



Estimating Daily Reproductive Number – Redistributed gap

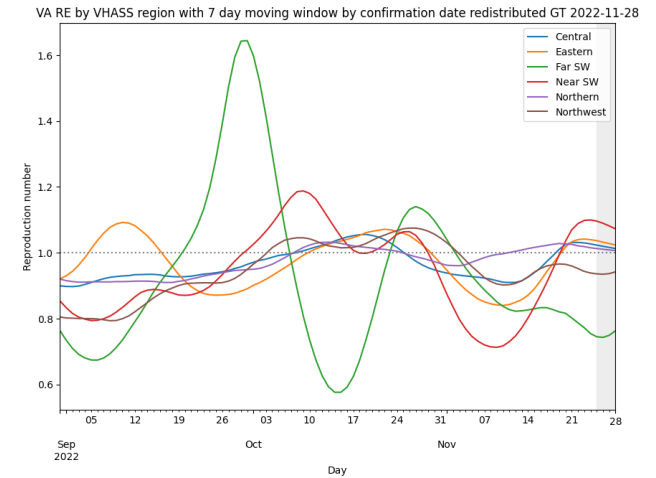
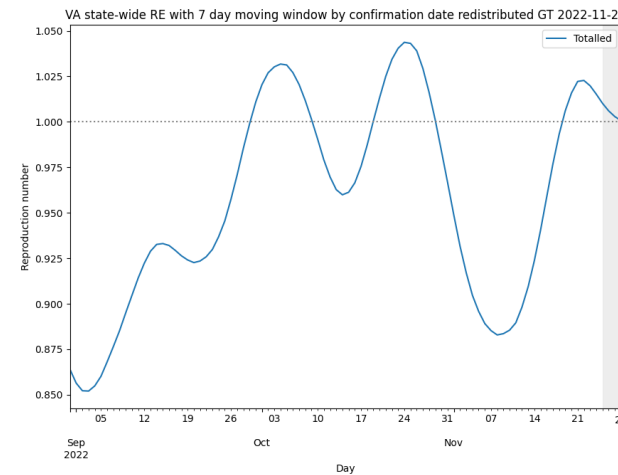
November 28th Estimates

Region	Date Confirmed R_e	Date Confirmed Diff Last Week
State-wide	1.001	0.137
Central	1.013	0.189
Eastern	1.024	0.211
Far SW	0.763	0.028
Near SW	1.073	0.329
Northern	1.007	0.046
Northwest	0.942	0.074

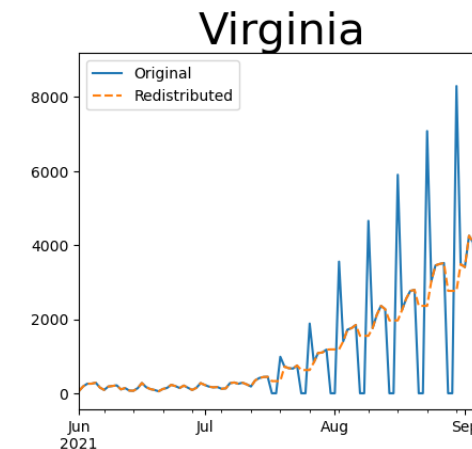
Methodology

- Wallinga-Teunis method (EpiEstim¹) for cases by confirmation date
- Serial interval: updated to discrete distribution from observations (mean=4.3, Flaxman et al, Nature 2020)
- Using Confirmation date since due to increasingly unstable estimates from onset date due to backfill

1. Anne Cori, Neil M. Ferguson, Christophe Fraser, Simon Cauchemez. A New Framework and Software to Estimate Time-Varying Reproduction Numbers During Epidemics. American Journal of Epidemiology, Volume 178, Issue 9, 1 November 2013, Pages 1505–1512, <https://doi.org/10.1093/aje/kwt133>



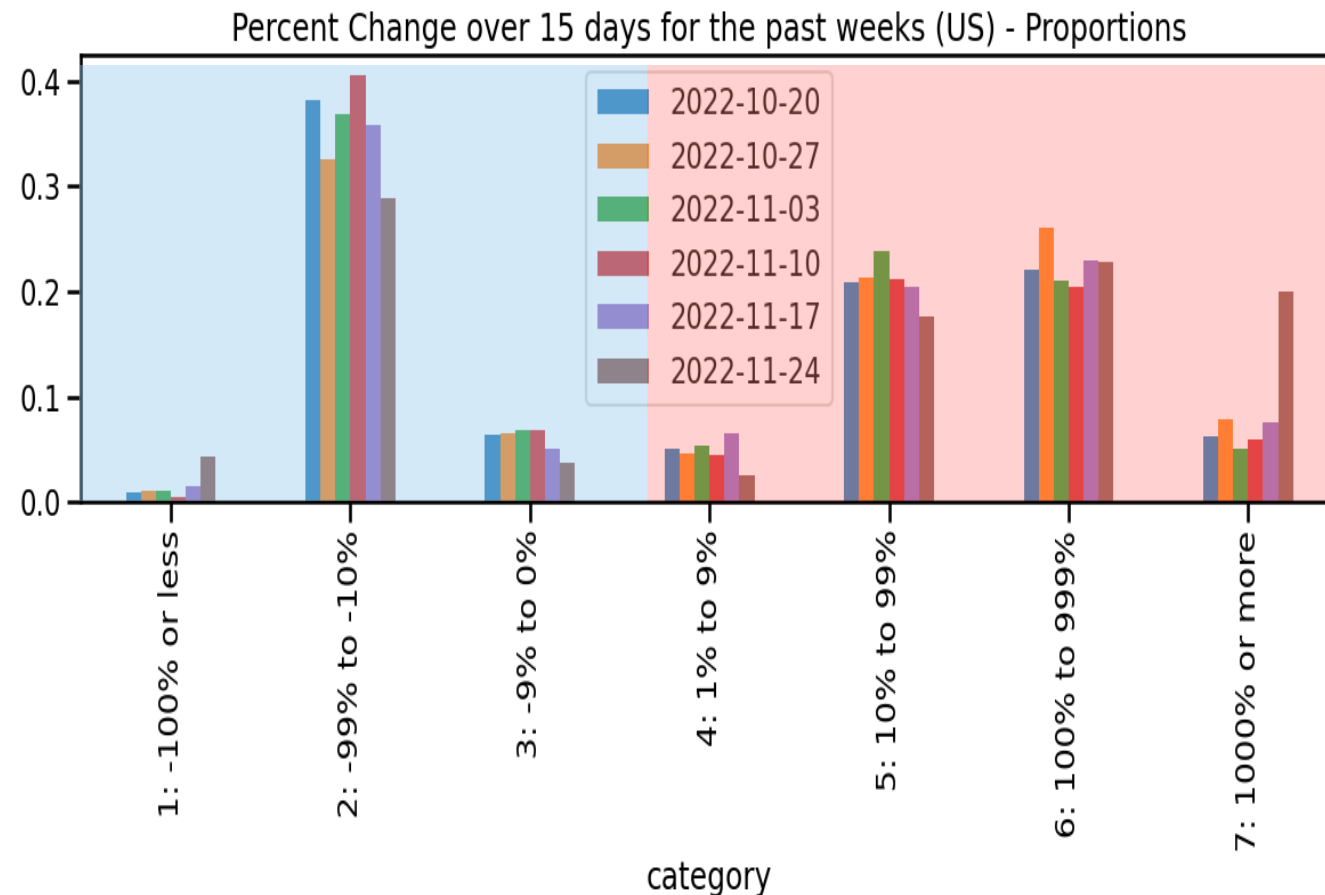
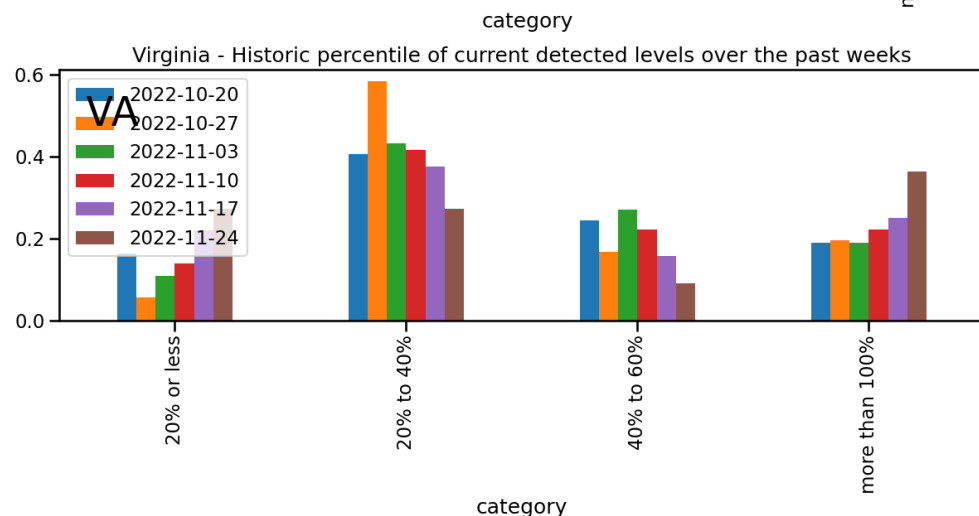
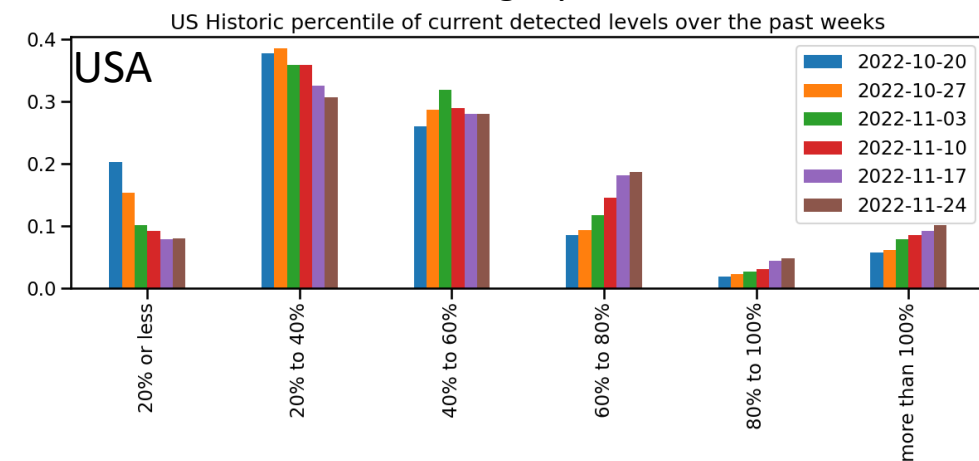
Skipping Weekend Reports & holidays biases estimates
Redistributed “big” report day to fill in gaps, and then estimate R from “smoothed” time series



Wastewater Monitoring

Wastewater provides a coarse early warning of COVID-19 levels in communities

- Overall in the US, there is an increase in sites with increased levels of virus compared to 15 days ago
- Growth seen in the category where current virus levels are at or exceeding max of previous historical levels

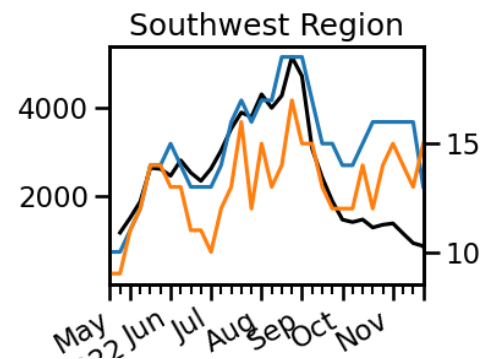
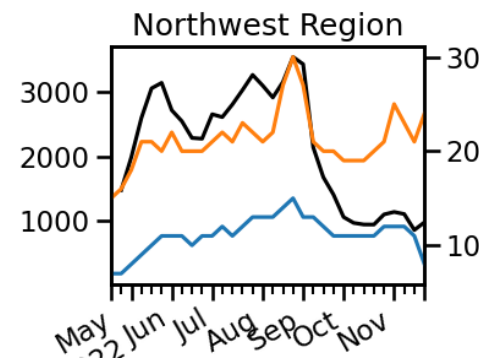
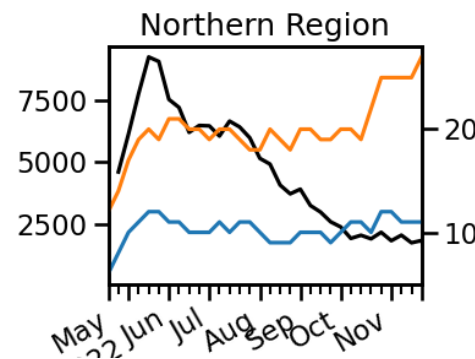
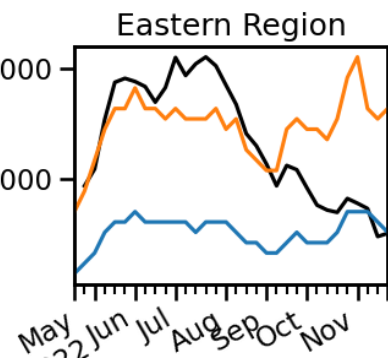
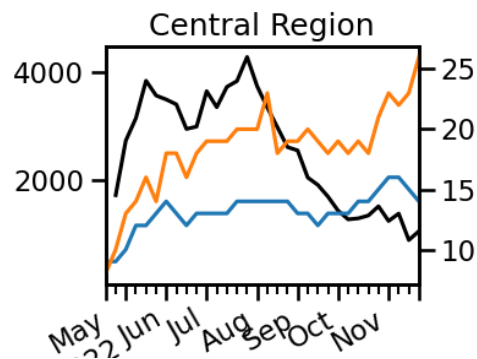
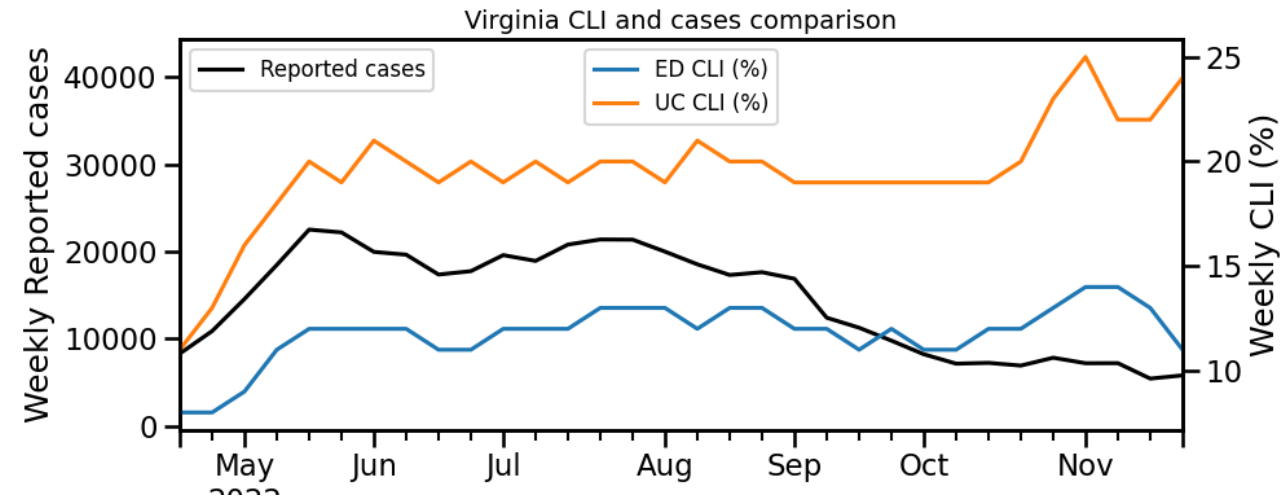


Data Source: [CDC Data Tracker](#)

COVID-like Illness Activity

COVID-like Illness (CLI) gives a measure of COVID transmission in the community

- Emergency Dept (ED) based CLI is more correlated with case reporting
- Urgent Care (UC) is a leading indicator but may be influenced by testing for other URIs
- **After 5 months of plateau, UC CLI remains higher than previous levels statewide**



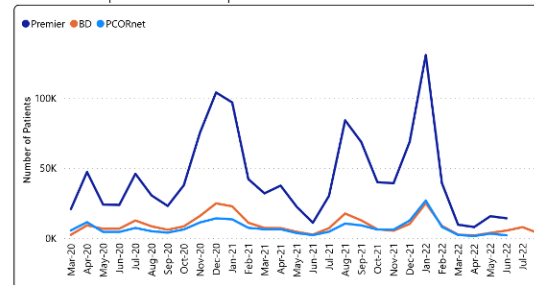
Hospitalizations and Severe Outcomes

Data Source: [CDC Data Tracker](#)

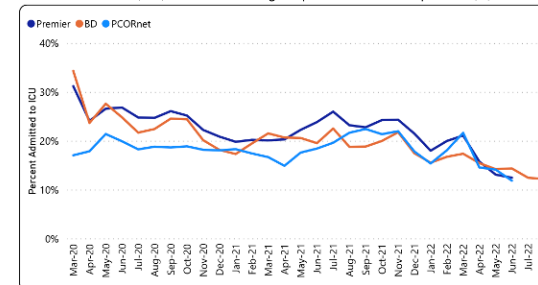
Proportion of most severe outcomes decreasing among those who are hospitalized

- ICU has declined from ~20% of hospitalized to 10-15% since initial Omicron wave
- Seen across all age-groups
- Recent rises in these rates have subsided in the past week

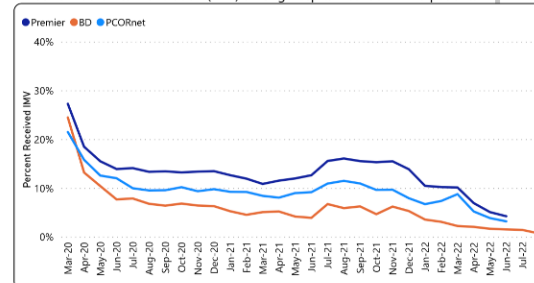
Number of hospitalized COVID-19 patients



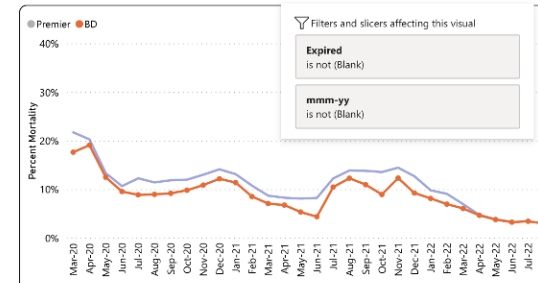
Intensive care unit (ICU) admission among hospitalized COVID-19 patients (%)



Invasive mechanical ventilation (IMV) among hospitalized COVID-19 patients

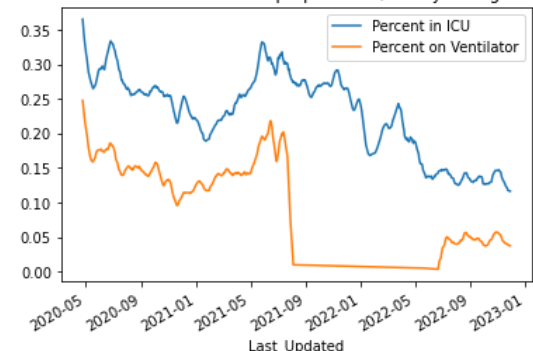


Mortality among hospitalized COVID-19 patients (%)



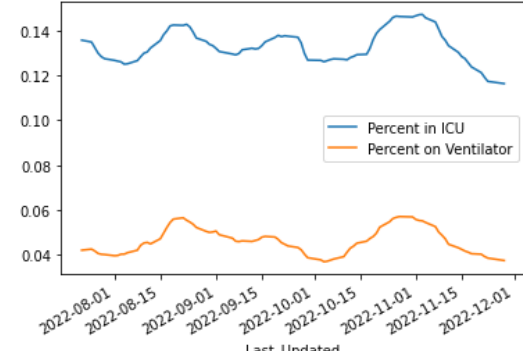
Virginia-wide – full pandemic

VA statewide ICU & Ventilation proportions (14 day rolling average)



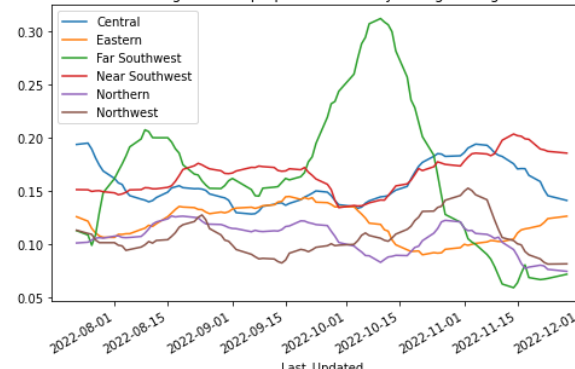
Virginia-wide – recent

VA statewide ICU & Ventilation proportions (14 day rolling average)



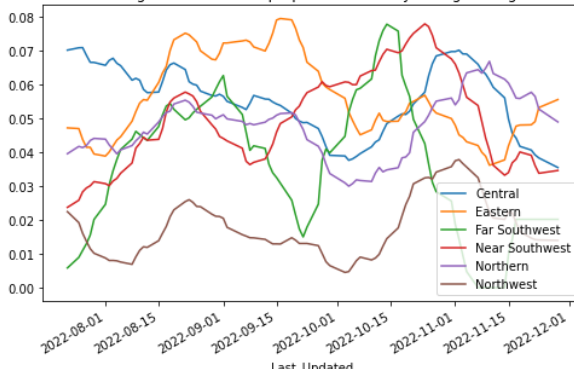
Virginia Regional ICU percent

VA Regional ICU proportions (14 day rolling average)



Virginia Regional Ventilation %

VA Regional Ventilation proportions (14 day rolling average)



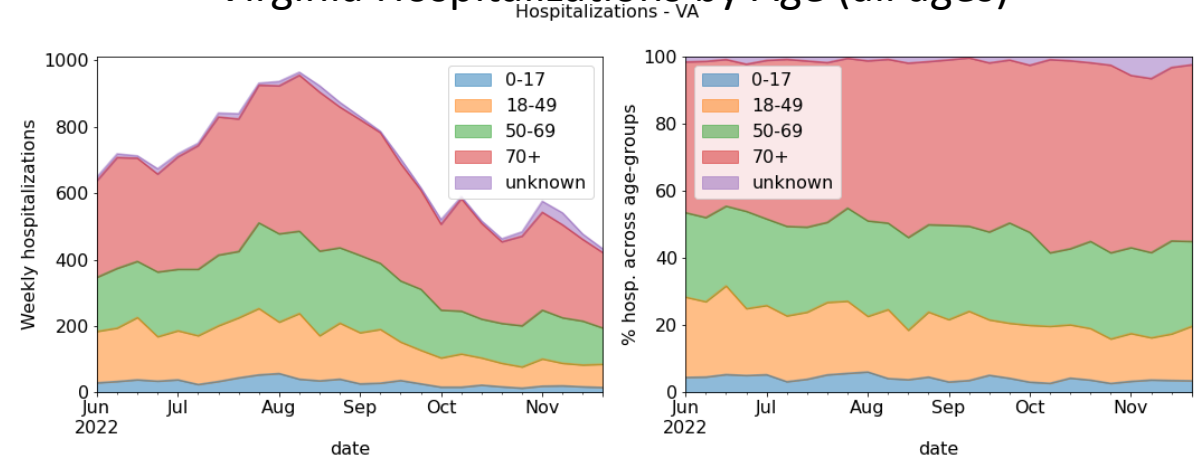
Hospitalizations in VA by Age

Age distribution in hospitals relatively stable

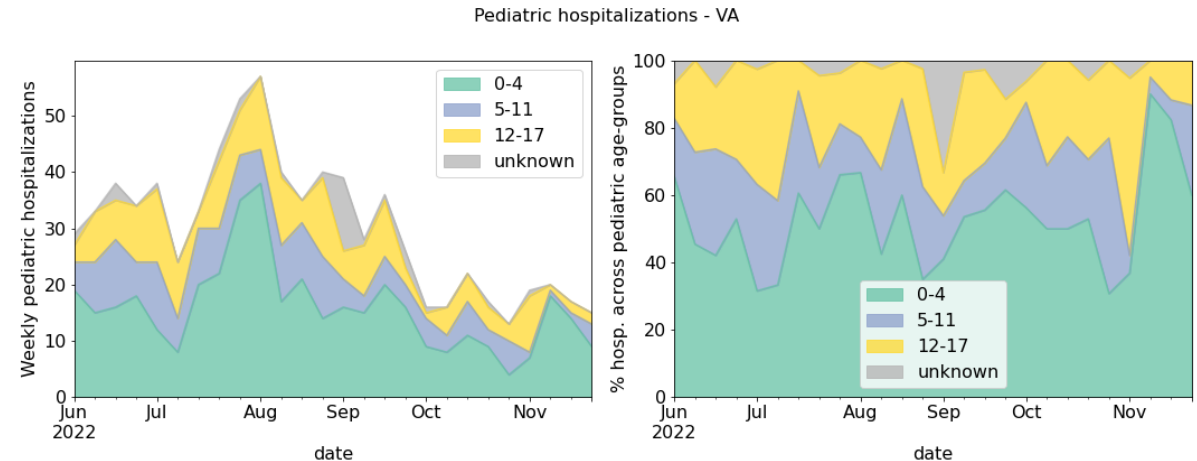
- Recent decline in overall pediatric hospitalizations
- Recent majority among pediatric hospitalizations has been in 0-4-yo

Note: These data are lagged and based on HHS hospital reporting

Virginia Hospitalizations by Age (all ages)



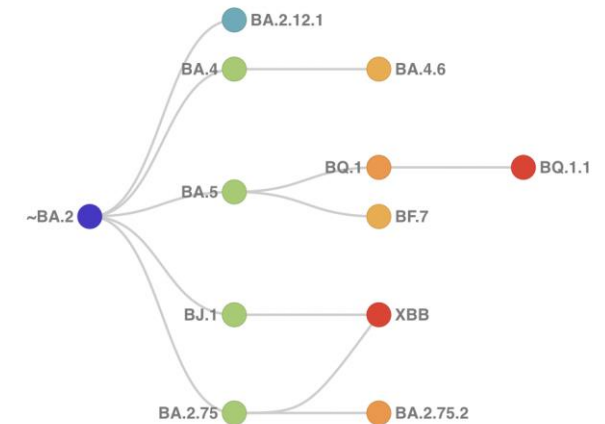
Pediatric Hospitalizations by Age (0-17yo)



SARS-CoV2 Variants of Concern

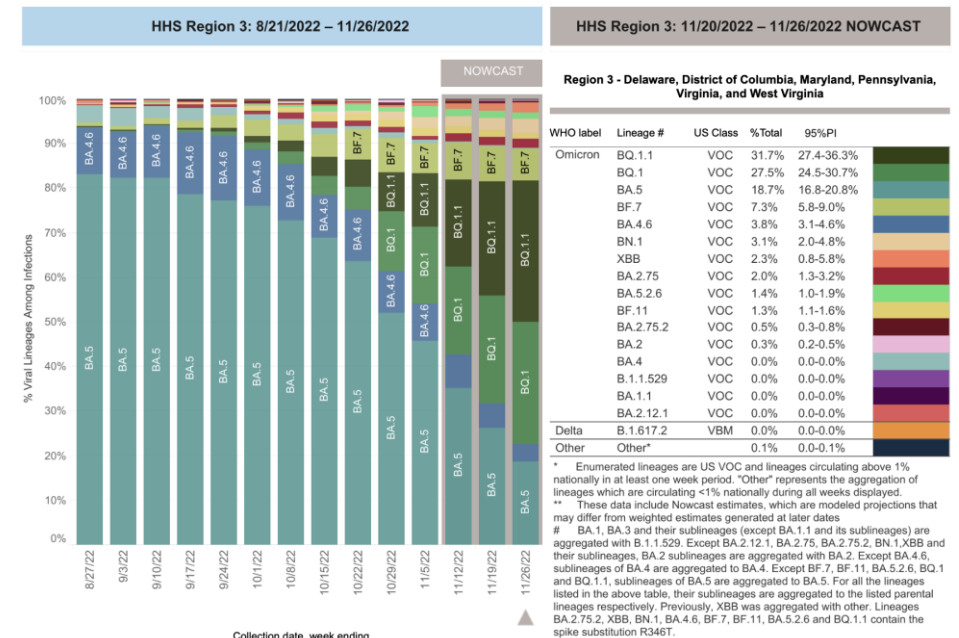
Emerging variants have potential to continue to alter the future trajectories of pandemic and have implications for future control

- **Variants have been observed to:** increase transmissibility, increase severity (more hospitalizations and/or deaths), and limit immunity provided by prior infection and vaccinations



Omicron Updates

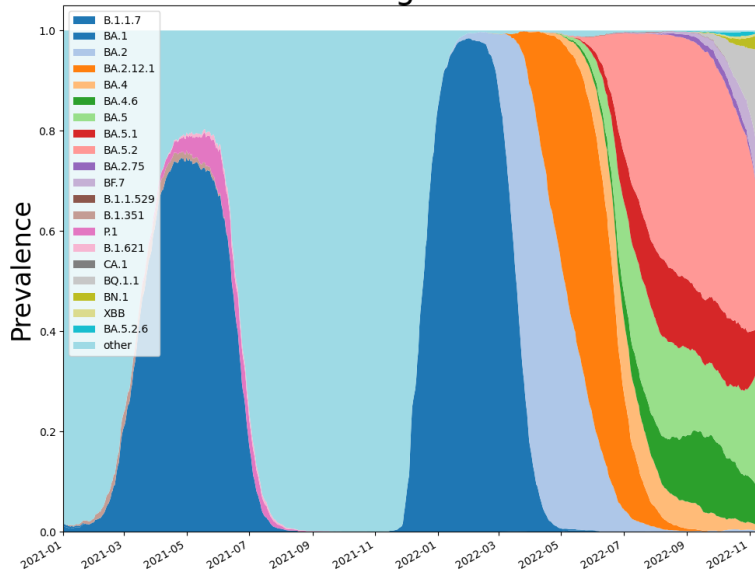
- Soup of tracked variants has grown to 82% from 74% last week
- BQ.1.1 remains at 32% with other BQ.1.* variants accounting for another 27%
- BA.2.75.* family variants, including BN.1, have shown more growth recently and now account for nearly a 6% share
- BF.7 and BA.4.6 have been slowly shrinking to 7% and 4%
- BA.5.2.6 and BN.1 are now broken out by CDC nowcast, and account for relatively smaller shares (3% and 4% respectively)
- XBB and subvariants remain a concern and now account for 2%



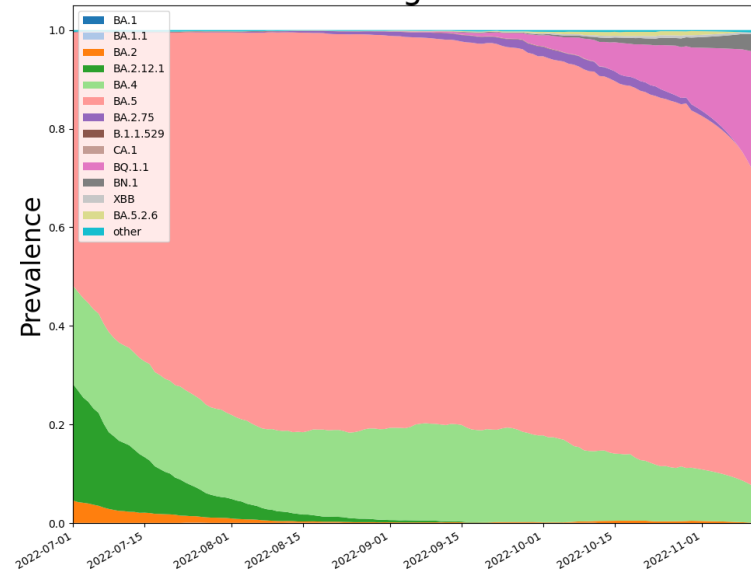
SARS-CoV2 Omicron Sub-Variants

As detected in whole Genomes in public repositories

Virginia

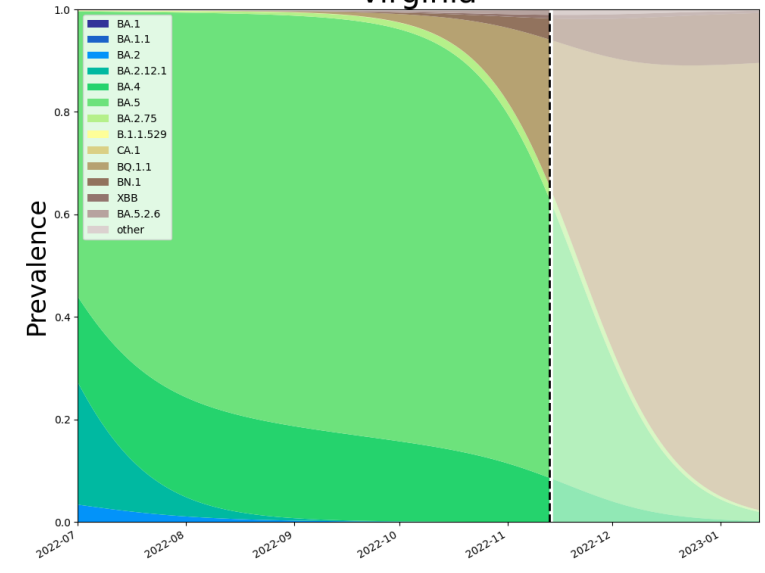


Virginia

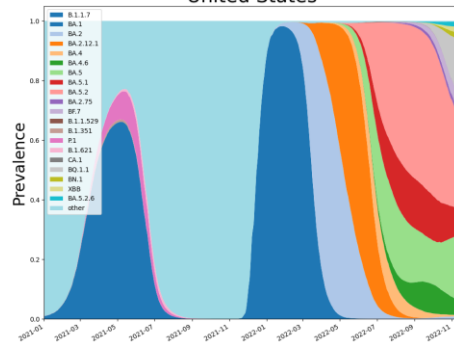


VoC Polynomial Fit Projections

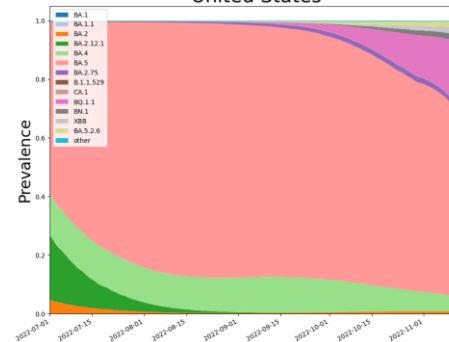
Virginia



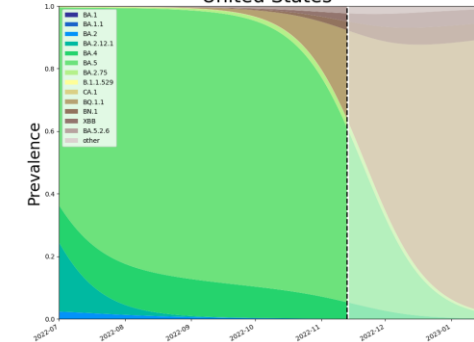
United States



United States



United States



Note: Data lags force projections to start in past. Everything from dotted line forward is a projection.

1-Dec-22

SARS-CoV2 Omicron Sub-Variants

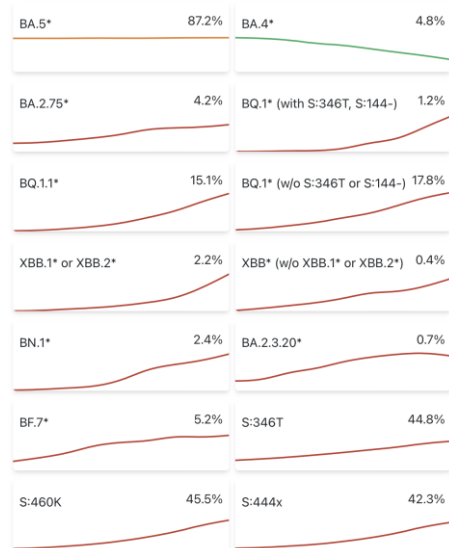
COV-spectrum

“Editor’s choice”
Variants to watch

Known variants

Which variant would you like to explore?

Editor's choice ▼

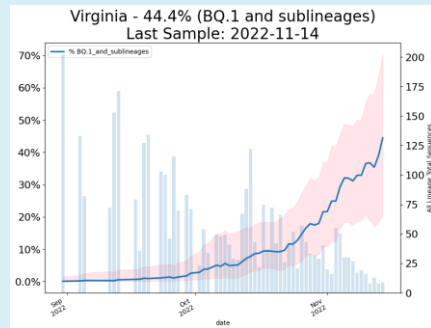
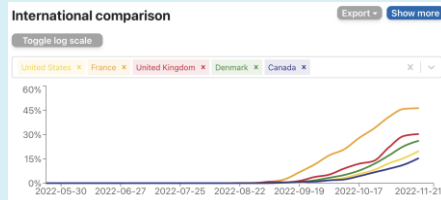
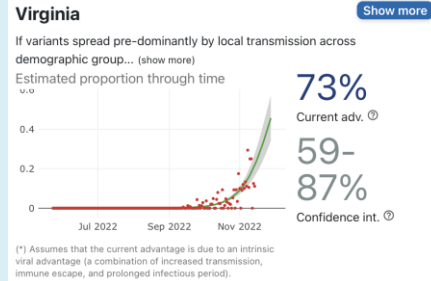


covSPECTRUM

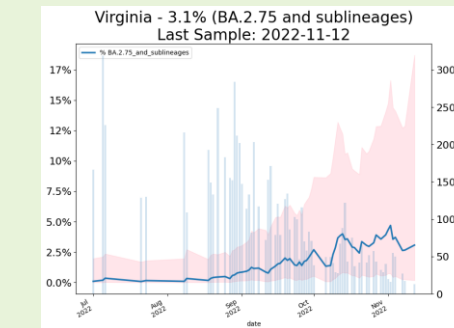
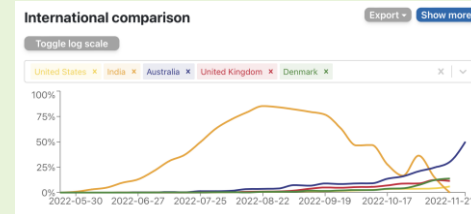
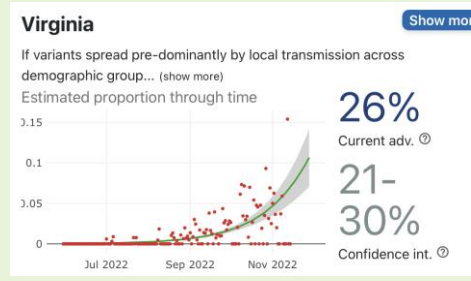
Enabled by data from **GISAI**D

1-Dec-22

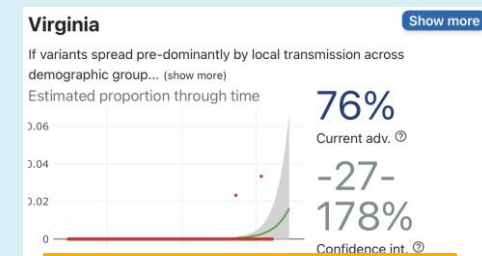
BQ.1.*



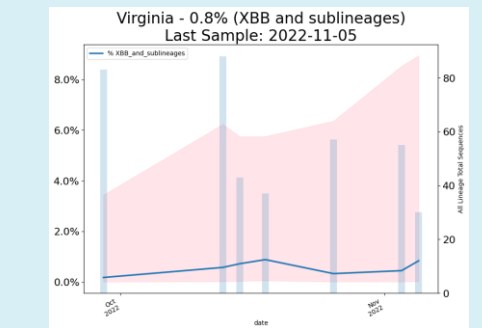
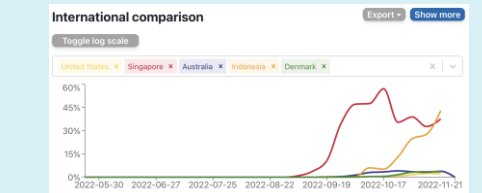
BA.2.75.*



XBB*



Limited samples, WIDE confidence bounds
Subject to large fluctuations

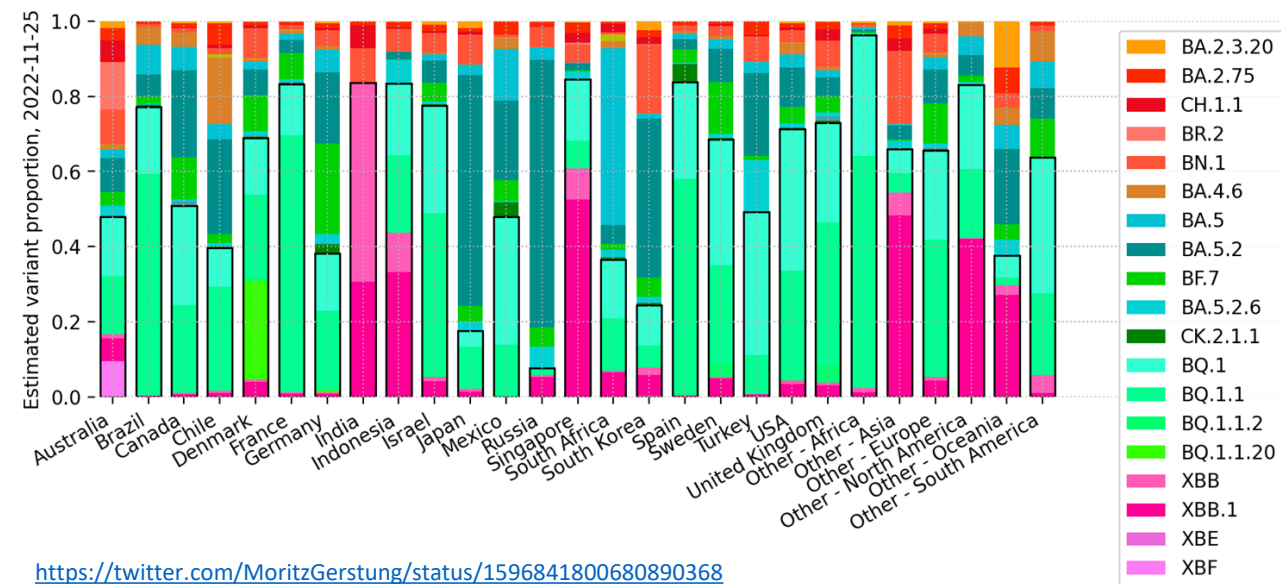
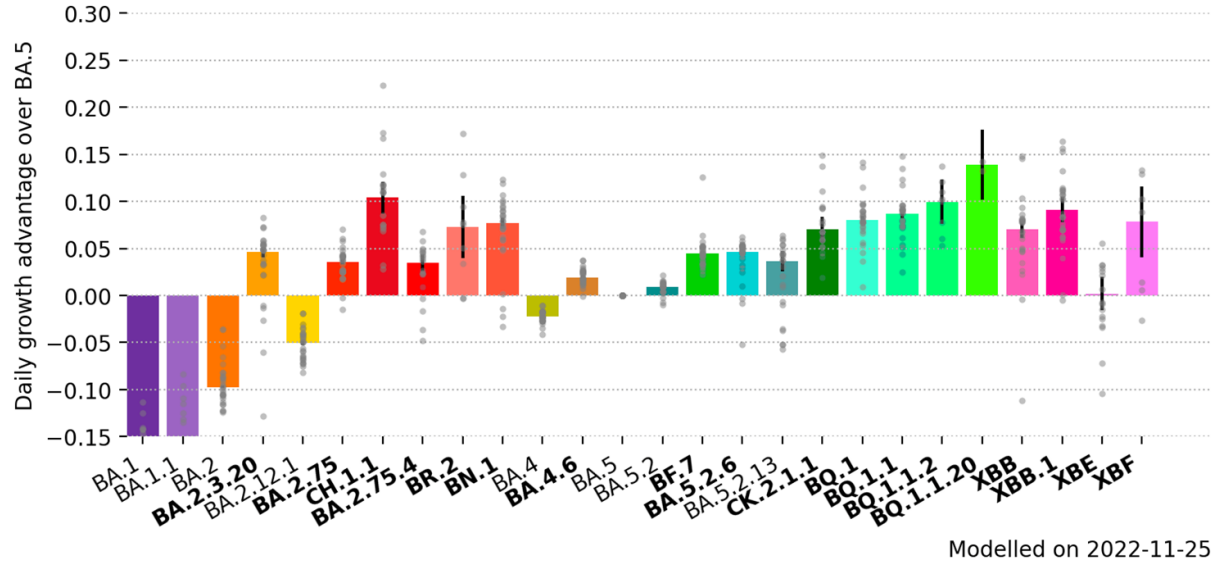
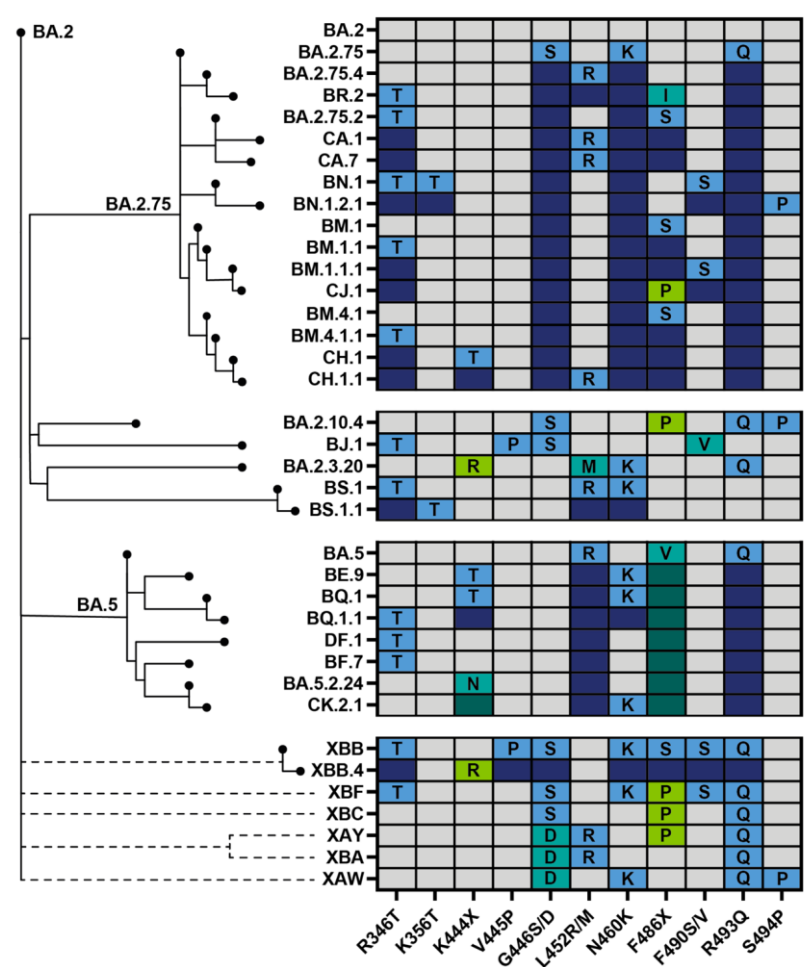


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Pandemic Pubs (Nov 29th, 2022)

1. SARS-CoV-2 surveillance yields both long phylogenetic branch lengths and ladder like mutational patterns, implicating both chronic infection and antigenic drift as likely contributors to fitness, resulting in multiple, potentially antigenically distinct lineages



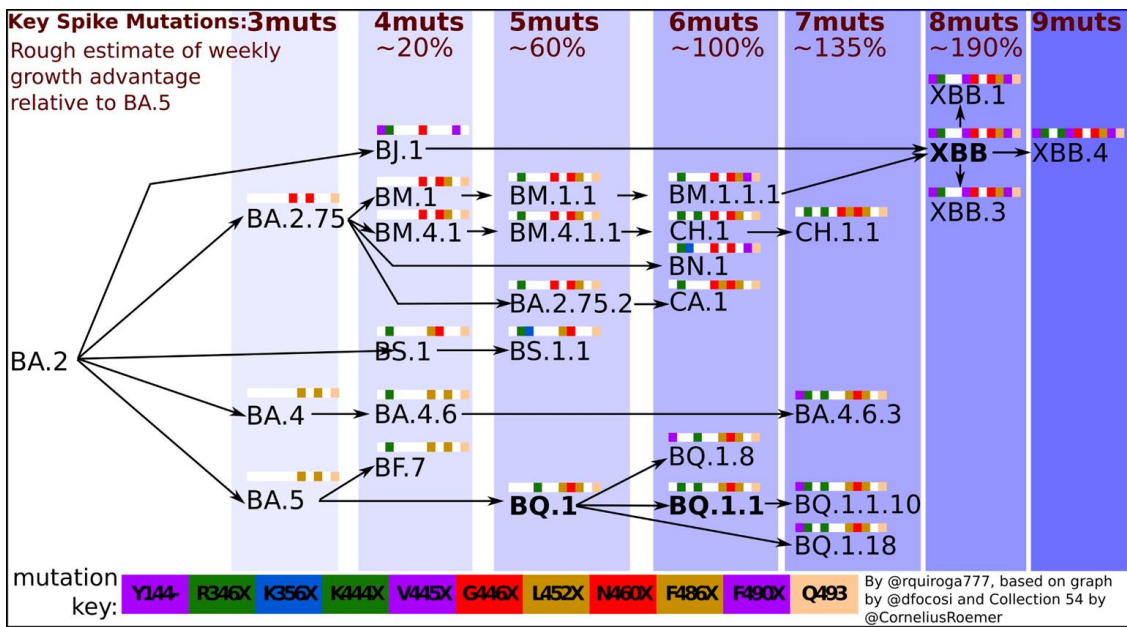
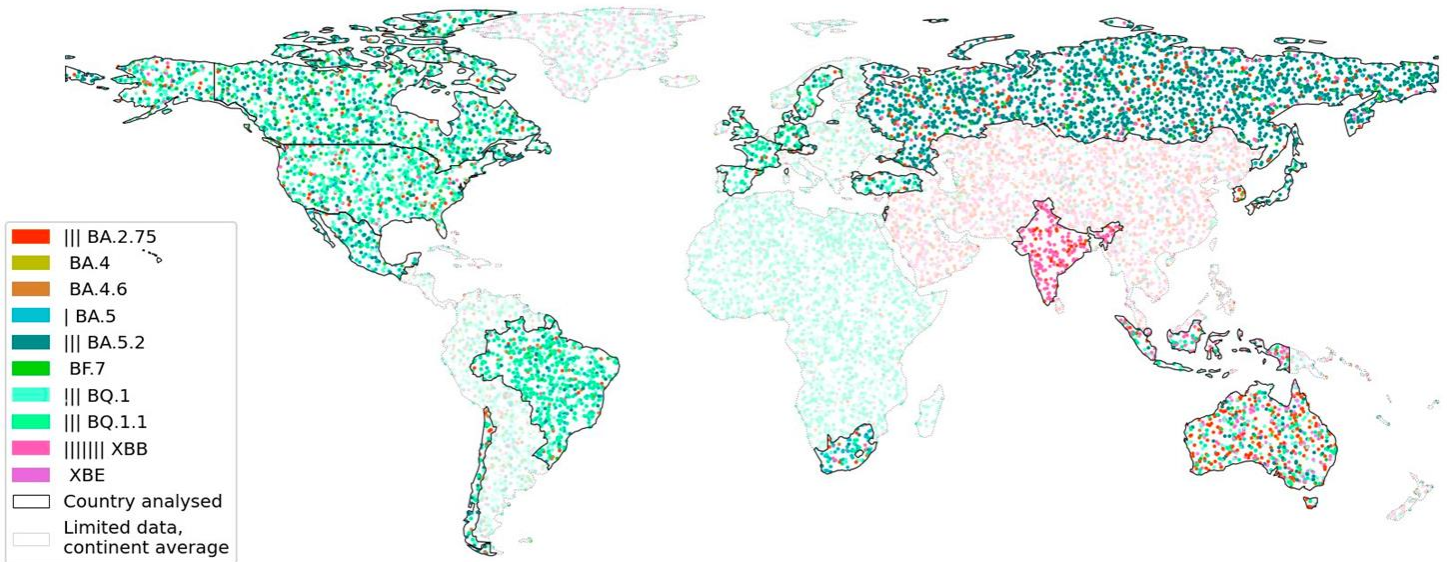
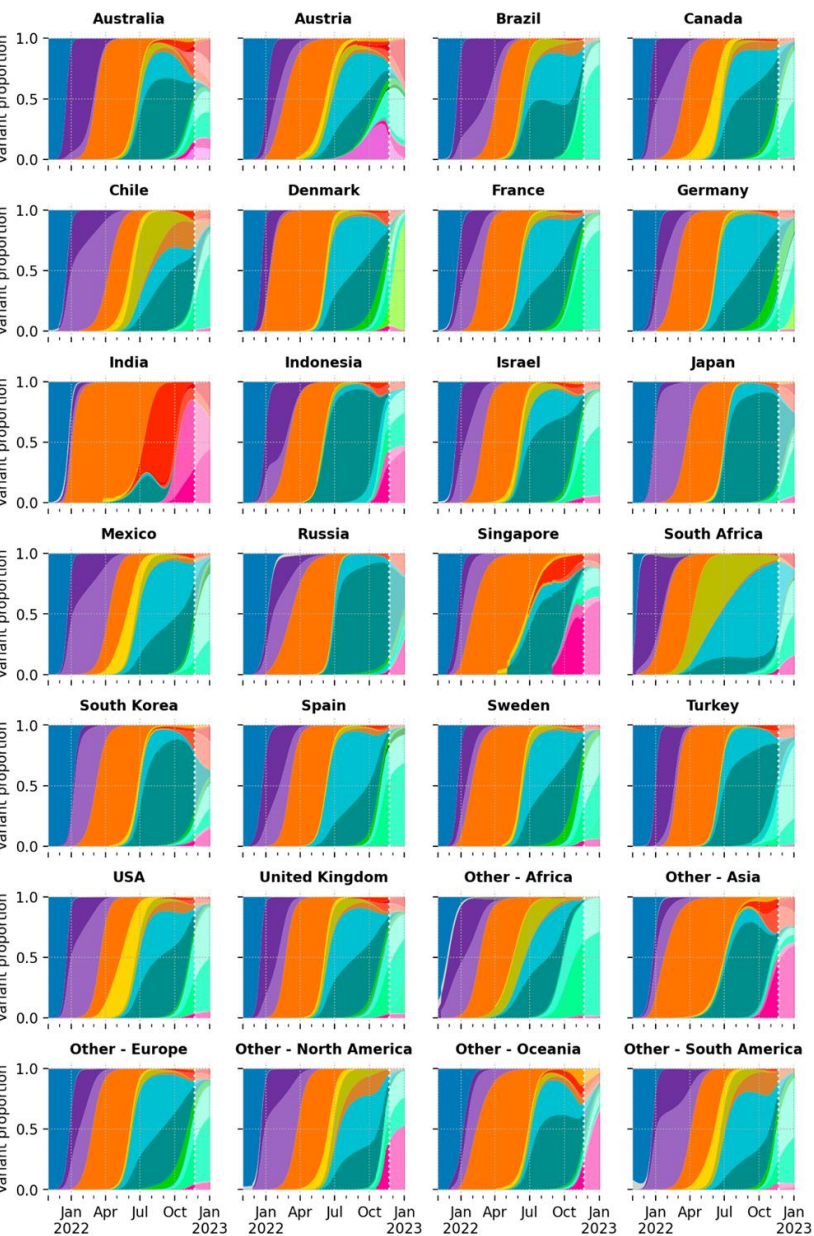
Authors from multiple countries highlight continued evidence that SARS-CoV-2 continues to diversify combinations of antigenically relevant mutations. **Authors raise the possibility that “several lineages could have similar enough growth rates, and enough antigenic distance from one another that they co-circulate, at least until a fitter lineage or variant emerges”.**

https://virological.org/t/sars-cov-2-evolution-post-omicron/911#post_1

<https://twitter.com/MoritzGerstung/status/1596841800680890368>

Pandemic Pubs (Nov 29th, 2022)

2. Variants around the world



<https://twitter.com/MoritzGerstung/status/1596841800680890368>
<https://twitter.com/PeacockFlu/status/1596492725171675136>

By @rquirola777, based on graph by @dfocosi and Collection 54 by @CorneliusRoemer

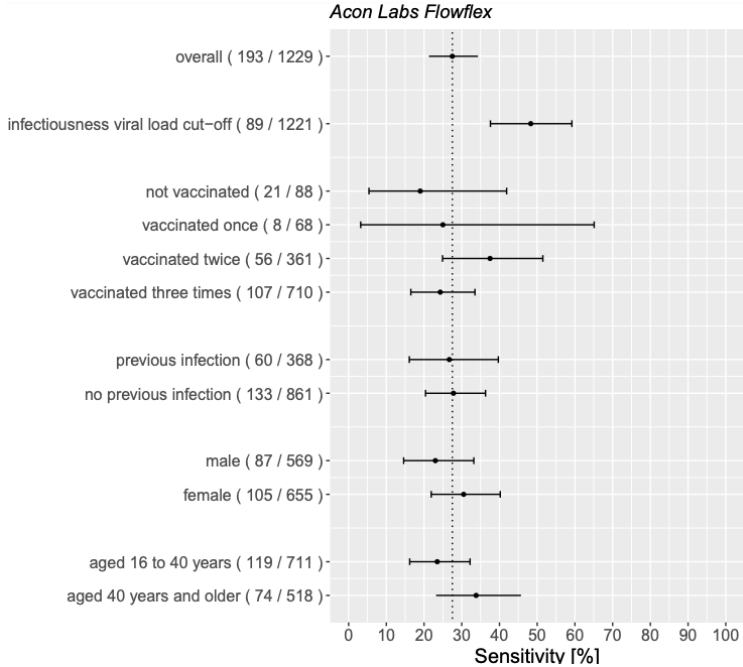
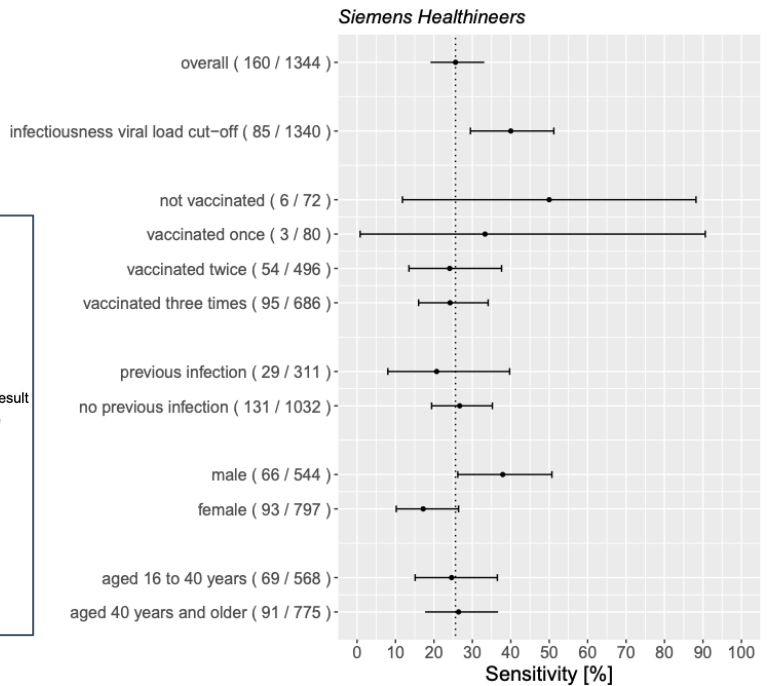
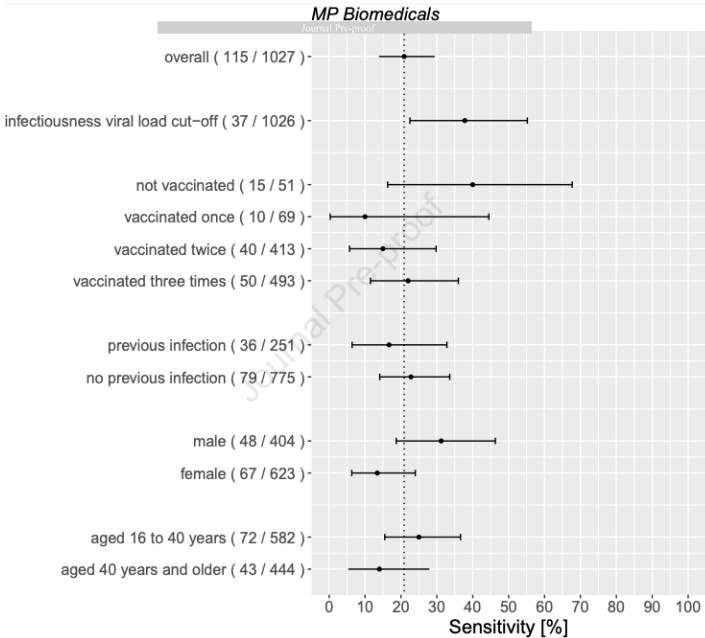
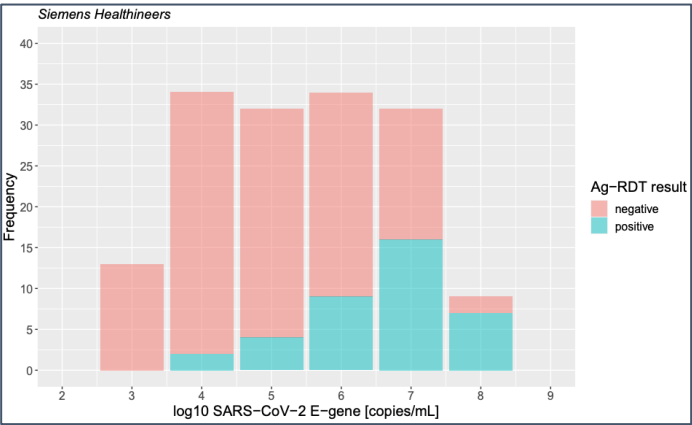
Pandemic Pubs (Nov 29th, 2022)

3. Netherlands study advocates, provides evidence for, repeated self-testing when asymptomatic testing to protect vulnerable individuals.

	N	RT-PCR test positivity* [%]	Sensitivity [%] (95%CI)	Specificity [%] (95%CI)	PPV [%] (95%CI)	NPV [%] (95%CI)
Flowflex						
Primary analysis	1229	15.7	27.5 (21.3-34.3)	99.8 (99.3-100)	96.4 (87.5-99.6)	88.1 (86.1-89.9)
Secondary (stratified) analyses:						
Viral load cut-off¶	1221	7.3	48.3 (37.6-59.2)	99.2 (98.5-99.6)	82.7 (69.7-91.8)	96.1 (94.8-97.1)
Vaccinated (at least once):						
Yes	1140	15.1	28.5 (21.9-35.9)	99.8 (99.3-100)	96.1 (86.5-99.5)	88.7 (86.7-90.5)
No	88	23.9	19.0 (5.4-41.9)	100 (94.6-100)	100 (39.8-100)	79.8 (69.6-87.7)
Previous SARS-CoV-2 infection:						
Yes	368	16.3	26.7 (16.1-39.7)	99.7 (98.2-100)	94.1 (71.3-99.9)	87.5 (83.5-90.7)
No	861	15.4	27.8 (20.4-36.3)	99.9 (99.2-100)	97.4 (86.2-99.9)	88.3 (85.9-90.4)
Sex:						
Female	655	16.0	30.5 (21.9-40.2)	99.8 (99.0-100)	97.0 (84.2-99.9)	88.3 (85.5-90.7)
Male	569	15.3	23.0 (14.6-33.2)	99.8 (98.8-100)	95.2 (76.2-99.9)	87.8 (84.7-90.4)
Age [years]:						
16-40	711	16.7	23.5 (16.2-32.2)	99.8 (99.1-100)	96.6 (82.2-99.9)	86.7 (83.9-89.1)
>40	518	14.3	33.8 (23.2-45.7)	99.8 (98.8-100)	96.2 (80.4-99.9)	90.0 (87.0-92.5)
MPBio						
Primary analysis	1027	11.2	20.9 (13.9-29.4)	99.8 (99.2-100)	92.3 (74.9-99.1)	90.9 (89.0-92.6)
Secondary (stratified) analyses:						
Viral load cut-off¶	1026	3.6	37.8 (22.5-55.2)	98.8 (97.9-99.4)	53.8 (33.4-73.4)	97.7 (96.6-98.5)
Vaccinated (at least once):						
Yes	975	10.3	18.0 (11.0-26.9)	99.8 (99.2-100)	90.0 (68.3-98.8)	91.4 (89.5-93.1)
No	51	29.4	40.0 (16.3-67.7)	100 (90.3-100)	100 (54.1-100)	80.0 (65.4-90.4)
Previous SARS-CoV-2 infection:						
Yes	251	14.3	16.7 (6.4-32.8)	100 (98.3-100)	100 (54.1-100)	87.8 (83.0-91.6)
No	775	10.2	22.8 (14.1-33.6)	99.7 (99.0-100)	90.0 (68.3-98.8)	91.9 (89.7-93.8)
Sex:						
Female	623	10.8	13.4 (6.3-24.0)	99.8 (99.0-100)	90.0 (55.5-99.7)	90.5 (87.9-92.7)
Male	404	11.9	31.2 (18.7-46.3)	99.7 (98.4-100)	93.8 (69.8-99.8)	91.5 (88.3-94.1)
Age [years]:						
16-40	582	12.4	25.0 (15.5-36.6)	99.8 (98.9-100)	94.7 (74.0-99.9)	90.4 (87.7-92.7)
>40	444	9.7	14.0 (5.3-27.9)	99.8 (98.6-100)	85.7 (42.1-99.6)	91.5 (88.5-94.0)
Clinitest						
Primary analysis	1344	11.9	25.6 (19.1-33.1)	99.9 (99.5-100)	97.6 (87.4-99.9)	90.9 (89.2-92.4)
Secondary (stratified) analyses:						
Viral load cut-off¶	1340	6.3	40.0 (29.5-51.2)	99.5 (99.0-99.8)	85.0 (70.2-94.3)	96.1 (94.9-97.1)
Vaccinated (at least once):						
Yes	1271	12.0	24.8 (18.2-32.5)	99.9 (99.5-100)	97.4 (86.5-99.9)	90.7 (88.9-92.2)
No	72	8.3	50.0 (11.8-88.2)	100 (94.6-100)	100 (29.2-100)	95.7 (87.8-99.1)
Previous SARS-CoV-2 infection:						
Yes	311	9.3	20.7 (8.0-39.7)	99.6 (98.0-100)	85.7 (42.1-99.6)	92.4 (88.9-95.1)
No	1032	12.7	26.7 (19.4-35.2)	100 (99.6-100)	90.4 (88.4-92.1)	90.4 (88.4-92.1)
Sex:						
Female	797	11.7	17.2 (10.2-26.4)	100 (99.5-100)	100 (79.4-100)	90.1 (87.8-92.1)
Male	544	12.1	37.9 (26.2-50.7)	99.8 (98.8-100)	96.2 (80.4-99.9)	92.1 (89.4-94.3)
Age [years]:						
16-40	568	12.1	24.6 (15.1-36.5)	99.8 (98.9-100)	94.4 (72.7-99.9)	90.5 (87.8-92.9)
>40	775	11.7	26.4 (17.7-36.7)	100 (99.5-100)	100 (85.8-100)	91.1 (88.8-93.0)

Researchers in the Netherlands performed a cross-sectional study in the Omicron period in three public health service covid-19 test sites in the **Netherlands including 3,600 asymptomatic individuals presenting for SARS-CoV-2 testing for any reason except confirmatory testing after a positive self-test. Sensitivities of three commonly used SARS-CoV-2 Ag-RDTs when used as self-tests in asymptomatic individuals in the Omicron period were very low.** Authors state “self-testing has limited value for asymptomatic individuals wishing to protect vulnerable persons and may even lead to a false sense of security.” Applying a viral load cut-off (≥5.2 log10 SARS-CoV-2 E-gene copies/mL), sensitivities increased to 48.3% (37.6 to 59.2%), 37.8% (22.5 to 55.2%), and 40.0% (29.5 to 51.2%), Acon Flowflex (Flowflex), MP Biomedicals (MPBio), and Siemens-Healthineers Clinitest (Clinitest) respectively. Cut-off established above which 95% of people with a positive RT-PCR test result had a positive virus culture in that previous study

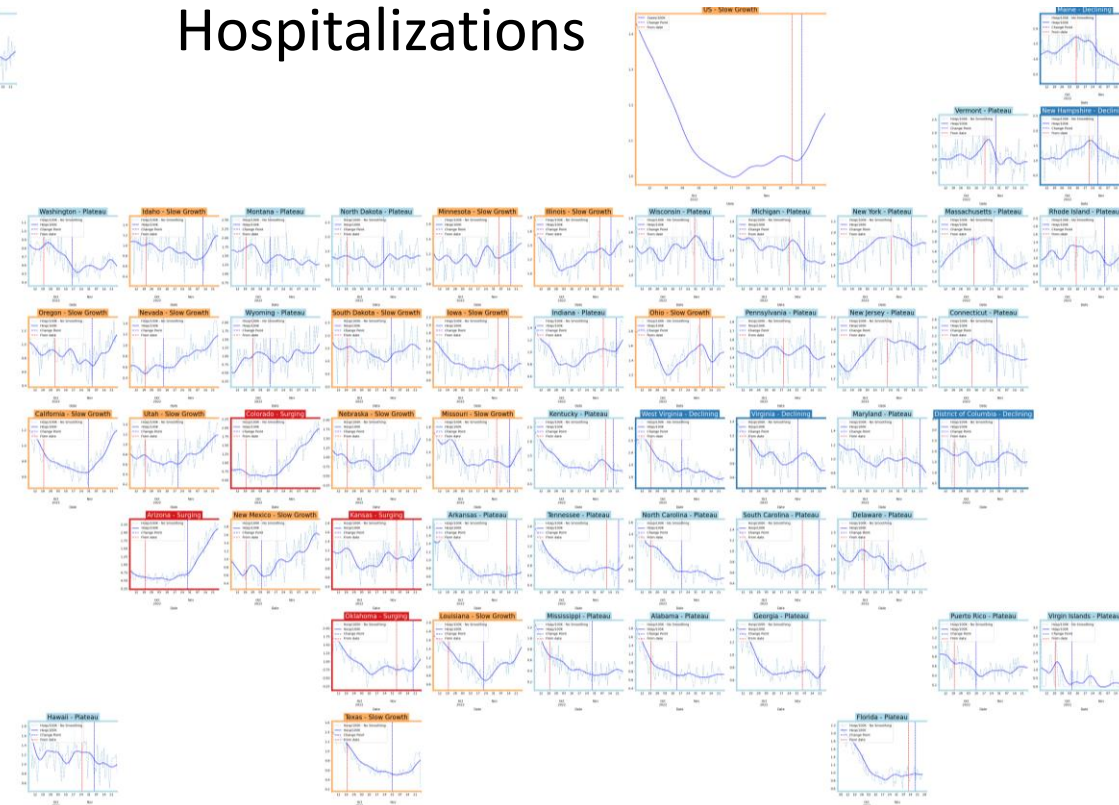
[https://www.clinicalmicrobiologyandinfection.com/article/S1198-743X\(22\)00570-5/fulltext#gr2](https://www.clinicalmicrobiologyandinfection.com/article/S1198-743X(22)00570-5/fulltext#gr2)



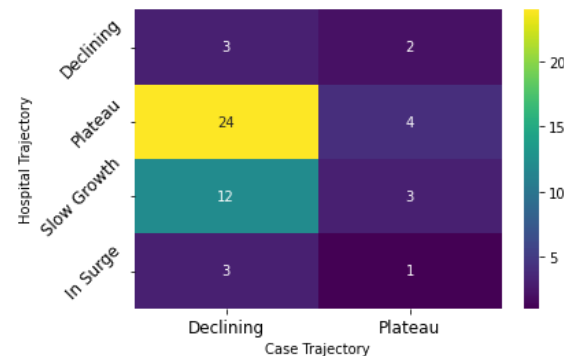
United States Cases & Hospitalizations



Hospitalizations



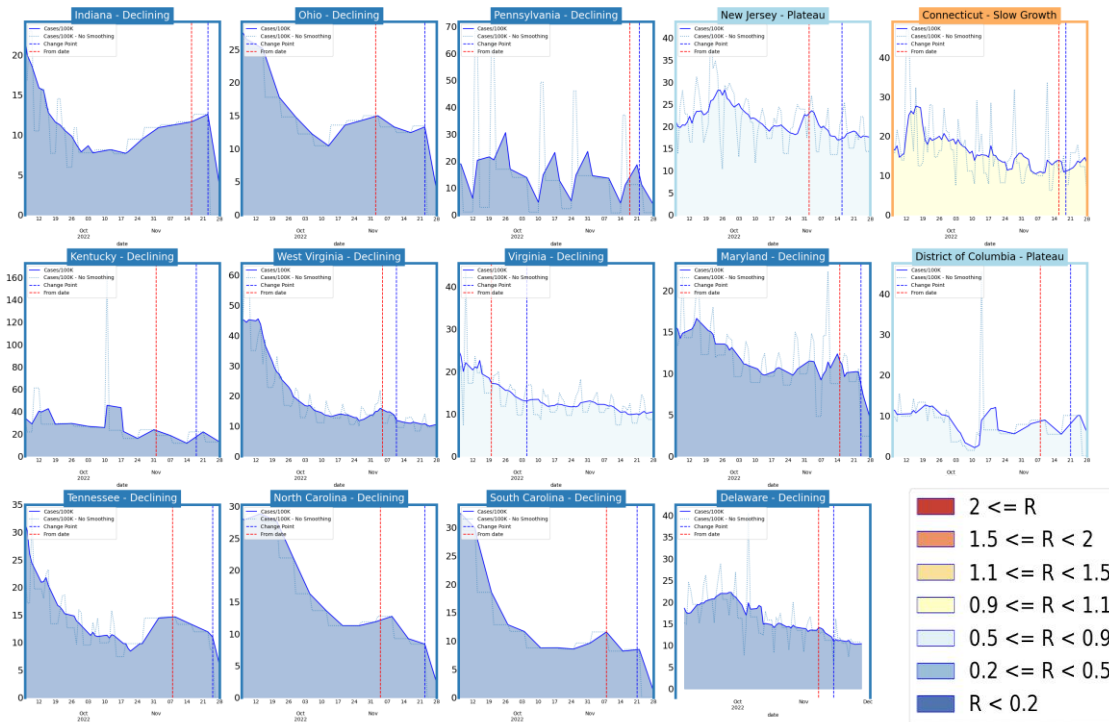
Status	Number of States	
	Current Week	Last Week
Declining	42	(40)
Plateau	11	(10)
Slow Growth	1	(2)
In Surge	0	(2)



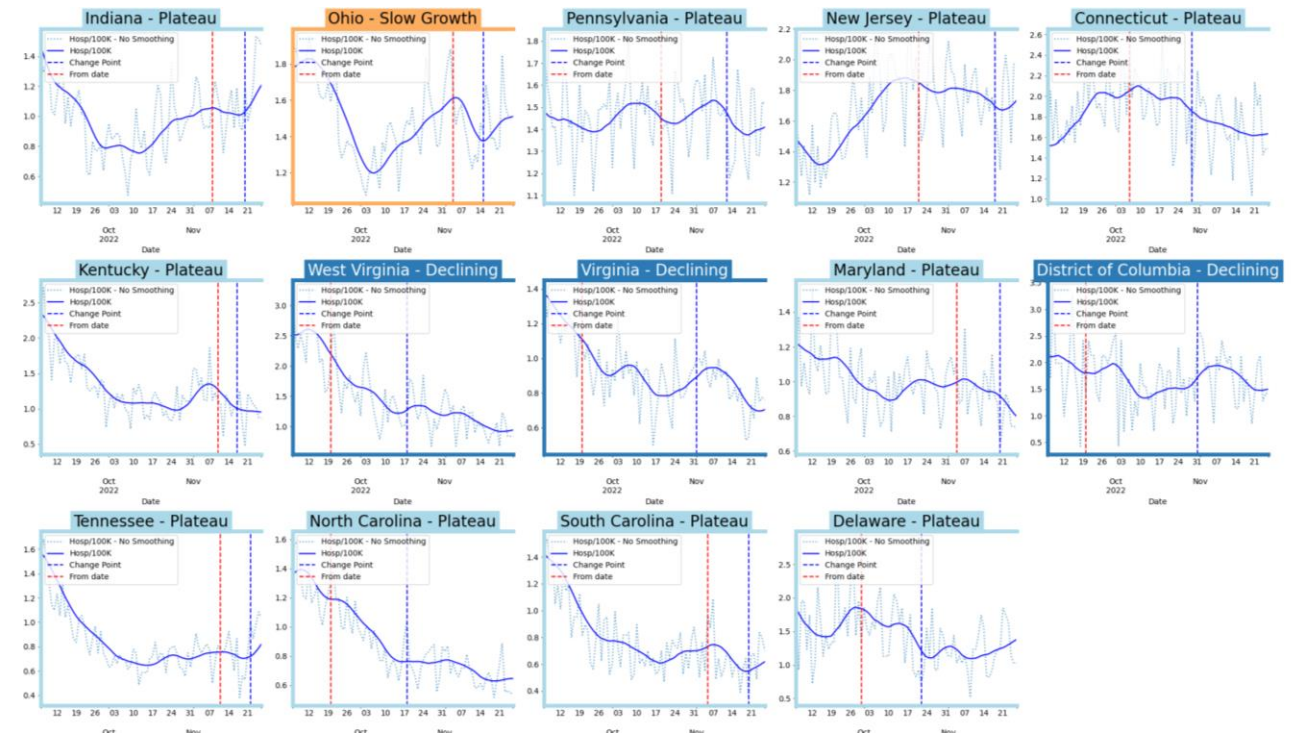
Status	Number of States	
	Current Week	Last Week
Declining	5	(4)
Plateau	29	(36)
Slow Growth	15	(11)
In Surge	4	(2)

Virginia and Her Neighbors

Cases

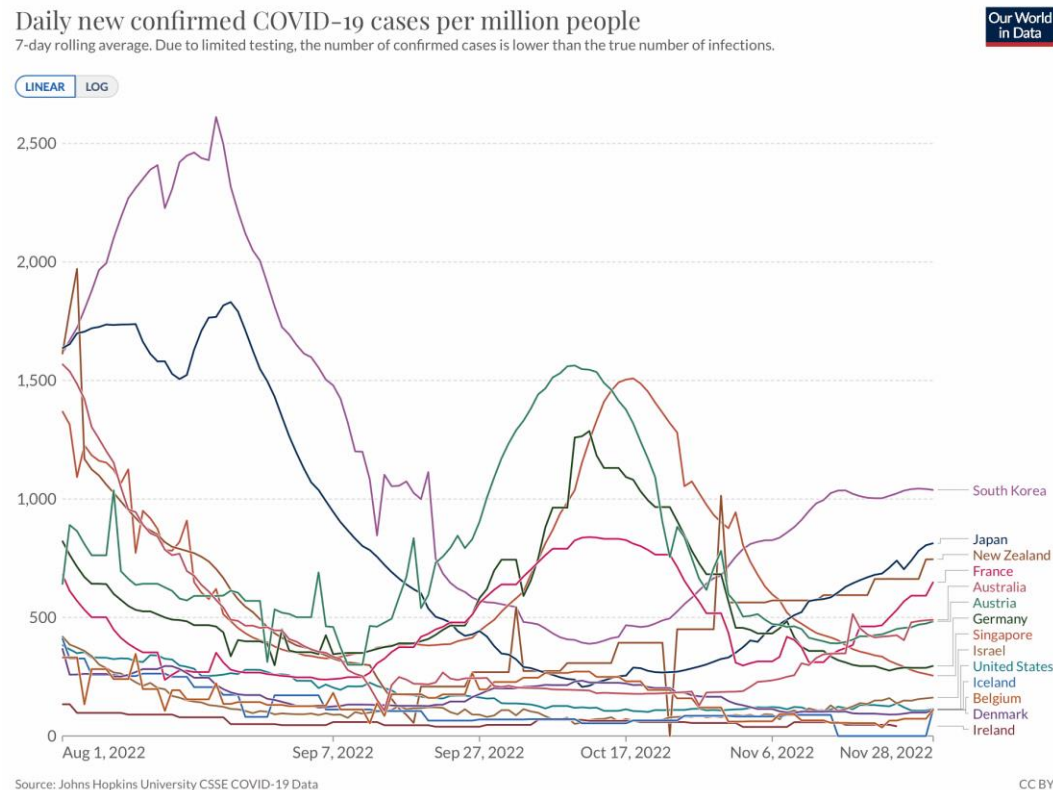


Hospitalizations

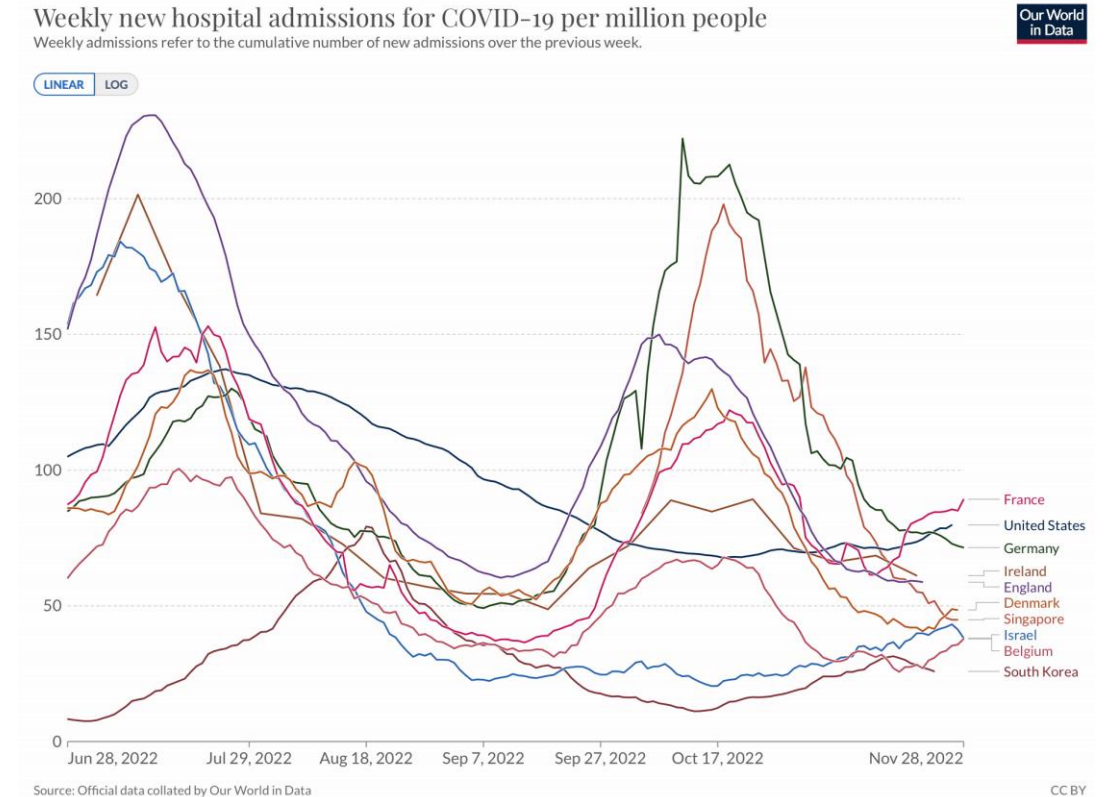


Around the World – Various trajectories

Confirmed cases



Hospitalizations



Zip code level weekly Case Rate (per 100K)

Case Rates in the last week by zip code

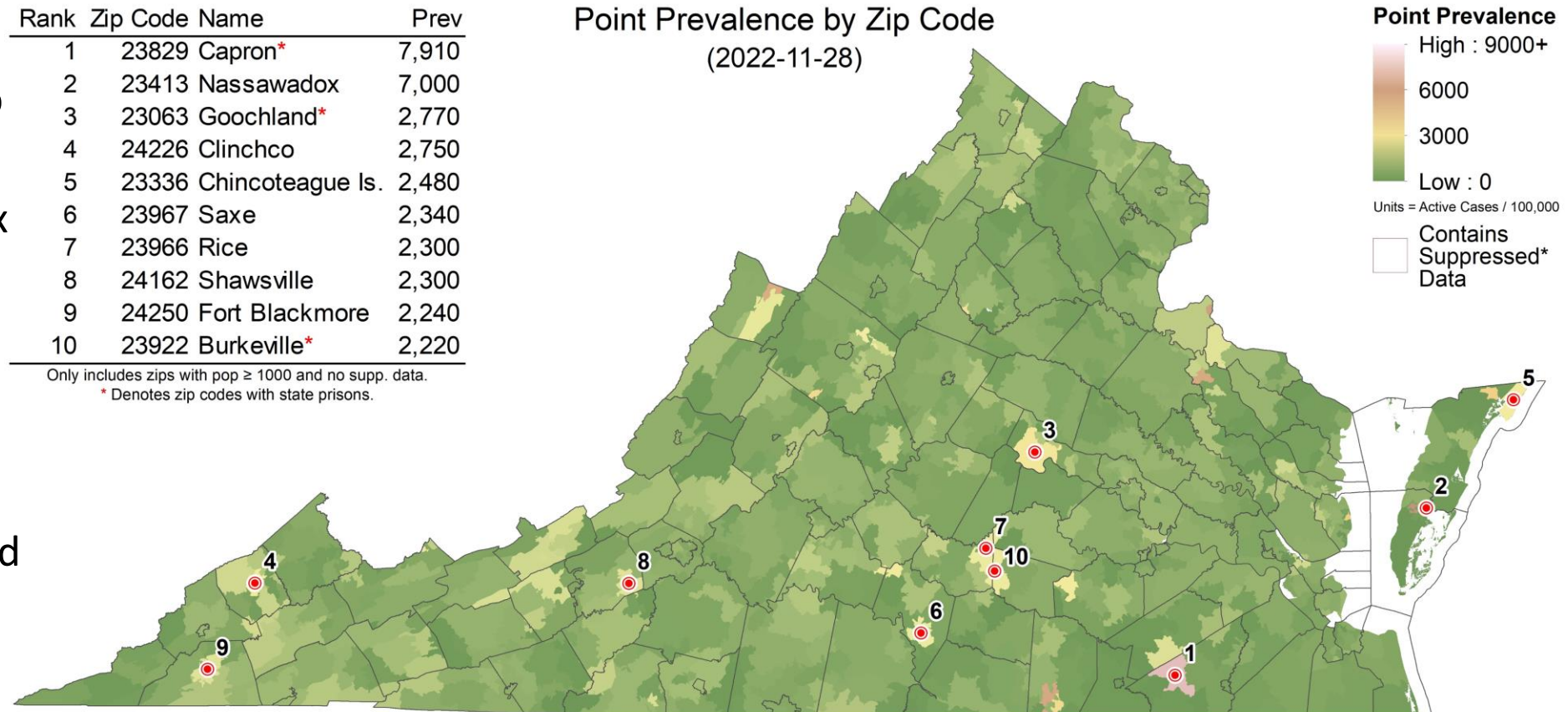
- Statewide rates remain quite low. We may need to adjust our color ramp for next week.
- Capron and Nassawadox are outliers with more than double any other zip's point prevalence.
- Three prisons appear in this week's top 10.
- Some counts are low and suppressed to protect anonymity. Those are shown with a dark red outline.

Rank	Zip Code	Name	Prev
1	23829	Capron*	7,910
2	23413	Nassawadox	7,000
3	23063	Goochland*	2,770
4	24226	Clinchco	2,750
5	23336	Chincoteague Is.	2,480
6	23967	Saxe	2,340
7	23966	Rice	2,300
8	24162	Shawsville	2,300
9	24250	Fort Blackmore	2,240
10	23922	Burkeville*	2,220

Only includes zips with pop ≥ 1000 and no supp. data.

* Denotes zip codes with state prisons.

Point Prevalence by Zip Code
(2022-11-28)

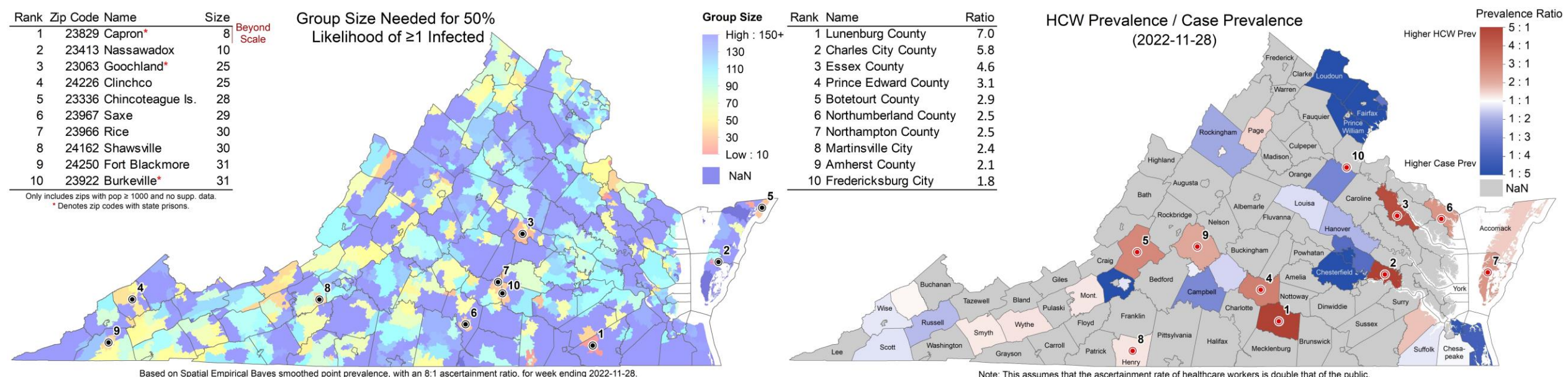


Based on Spatial Empirical Bayes smoothed point prevalence, with an 8:1 ascertainment ratio, for week ending 2022-11-28.

Risk of Exposure by Group Size and HCW prevalence

Case Prevalence in the last week by zip code used to calculate risk of encountering someone infected in a gathering of randomly selected people

- **Group Size:** Assumes **8 undetected infections** per confirmed case (ascertainment rate from recent seroprevalence survey) and shows minimum size of a group with a 50% chance an individual is infected by zip code (e.g., in a group of 8 in Capron, there is a 50% chance someone will be infected).
- **HCW ratio:** Case rate among health care workers (HCW) in the last week using patient facing health care workers as the denominator / population's case prevalence. Madison Co. is no longer in the top 10.



Current Hot-Spots

Case rates that are significantly different from neighboring areas or model projections

- **Spatial:** Getis-Ord Gi* based hot spots compare clusters of zip codes with weekly case prevalence higher than nearby zip codes to identify larger areas with statistically significant deviations
- **Temporal:** The weekly case rate (per 100K) projected last week compared to observed by county, which highlights temporal fluctuations that differ from the model's projections.
- Capron and Nassawadox lead the spatial hotspots. Temporal hotspots are primarily found in central Virginia and along the coast. Model forecasts tracked cases very well, and there was no residual autocorrelation.

Spatial Hotspots

Spot	Zip Code	Name	Conf.
1	23829	Capron*	99%
2	23413	Nassawadox	99%
3	23063	Goochland*	95%

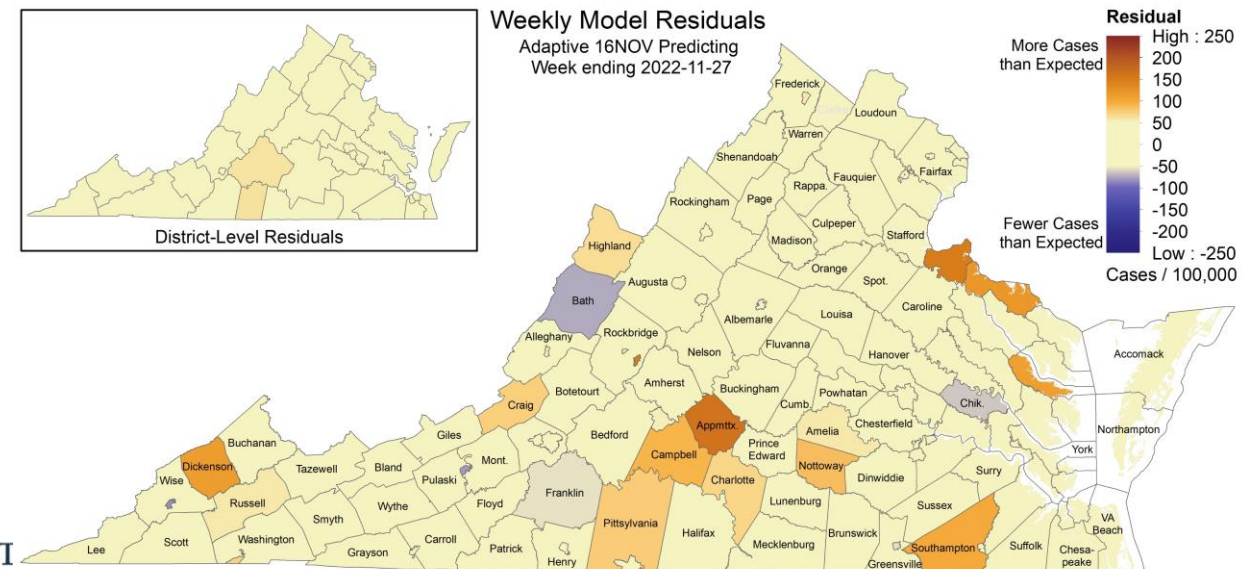
Only zips with pop ≥ 1000 and no supp. data.
* Denotes zip codes with state prisons.

Point Prevalence Hot Spots by Zip Code
(2022-11-28)



Based on Global Empirical Bayes smoothed point prevalence for week ending 2022-11-28.

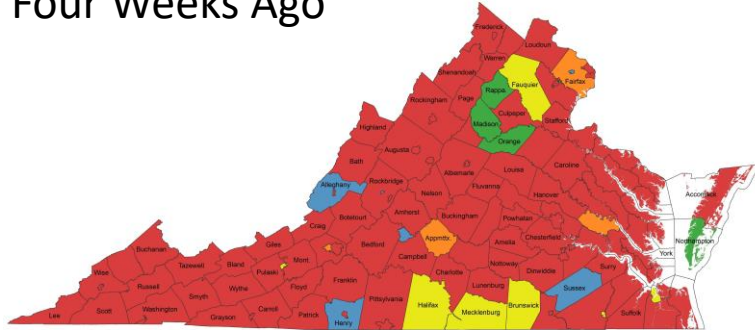
Clustered Temporal Hotspots



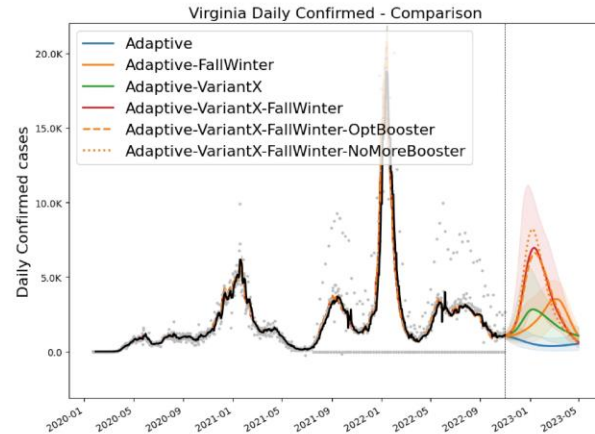
Scenario Trajectory Tracking

Which scenario from a month ago did projection for each county track closest?

Four Weeks Ago

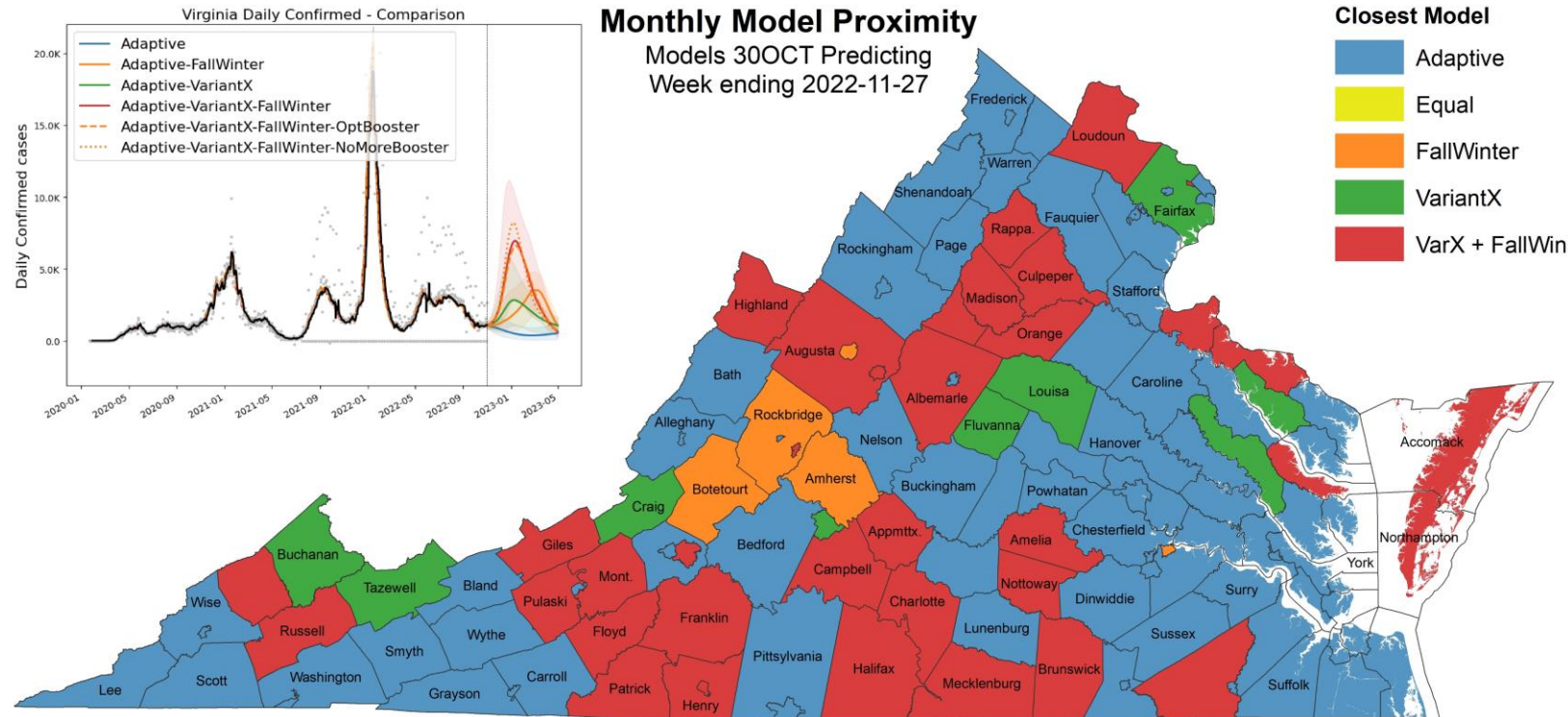


Two Weeks Ago



Monthly Model Proximity

Models 30OCT Predicting
Week ending 2022-11-27



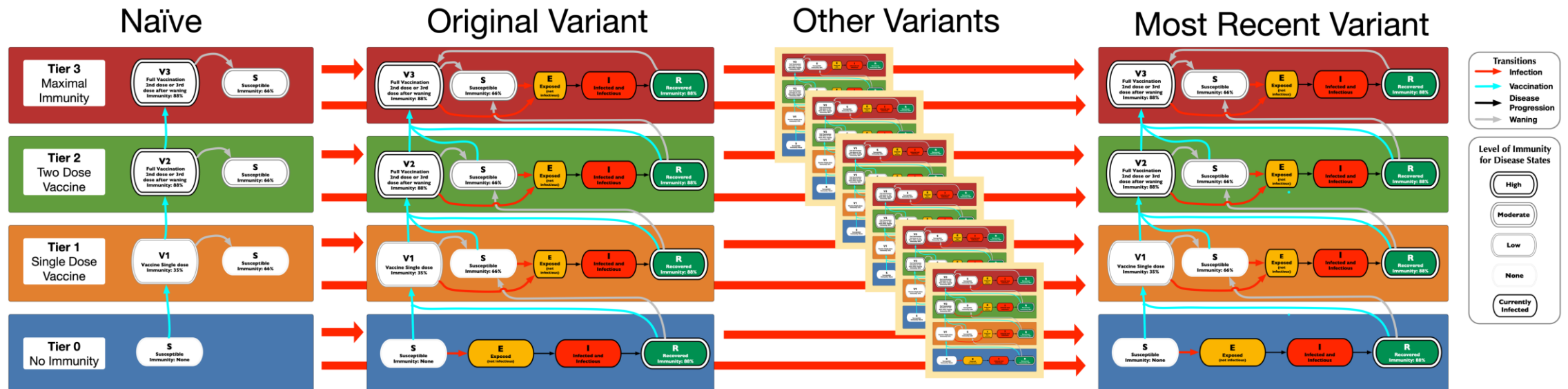
- One-month projections separate the scenarios more clearly and reveal larger overall patterns.
- Over the last six weeks, the Adaptive scenario forecasts have started to outperform those of FallWinter-variantX in an increasing number of counties. Adaptive was most accurate in 82 of 133 locales this week.

Model Update – Adaptive Fitting

Model Structure Extended for more sub-variants

Omicron sub-variants escape immunity induced by previous sub-variants

- Multiple strain support allows representation of differential protection based on immunological history (BA.1, BA.2, BA.2.12.1, BA.4/5, and future variants (VariantX))
- Each sub-variant has differing levels of immune escape to previous sub-variants, the prevalences are based on observations for fitting purposes, and projections use estimated future prevalences
- Adaptive fitting approach continues to use simulation to generate the full distribution of immune states across the population



Adaptive Fitting Approach

Each county fit precisely, with recent trends used for future projection

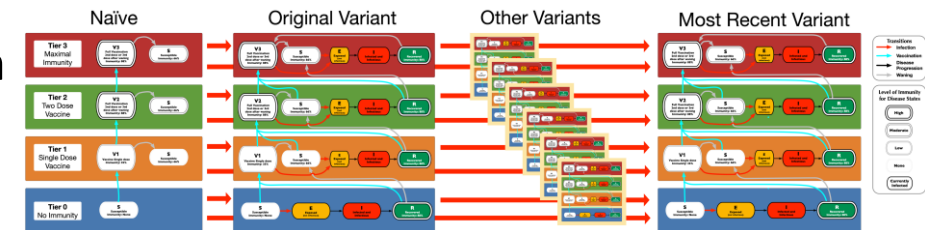
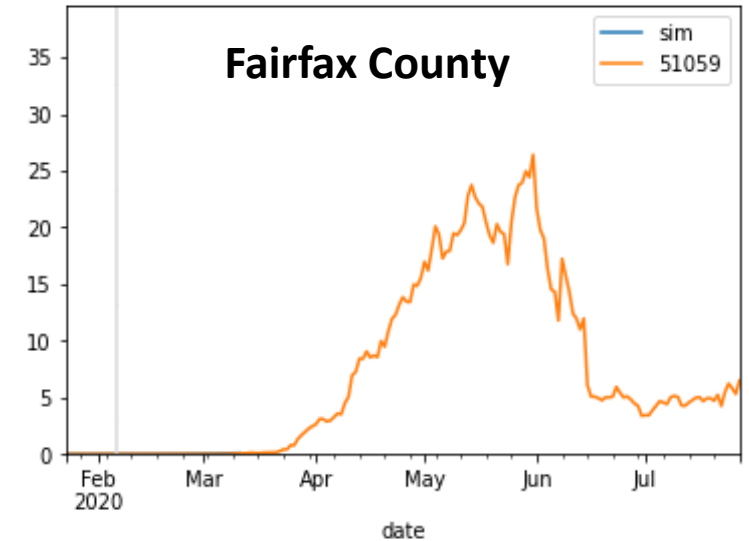
- Allows history to be precisely captured, and used to guide bounds on projections

Model: An alternative use of the same meta-population model, PatchSim with multiple tiers of immunity

- Allows for future “what-if” Scenarios to be layered on top of calibrated model
- Allows for waning of immunity and for partial immunity against different outcomes (eg lower protection for infection than death)

External Seeding: Steady low-level importation

- Widespread pandemic eliminates sensitivity to initial conditions, we use steady 1 case per 10M population per day external seeding



Calibration Approach

- **Data:**
 - County level case counts by date of onset (from VDH)
 - Confirmed cases for model fitting
- **Calibration:** fit model to observed data and ensemble's forecast
 - Tune transmissibility across ranges of:
 - Duration of incubation (5-9 days), infectiousness (3-7 days)
 - Undocumented case rate (1x to 7x) guided by seroprevalence studies
 - Detection delay: exposure to confirmation (4-12 days)
 - Approach captures uncertainty, but allows model to precisely track the full trajectory of the outbreak
- **Project:** future cases and outcomes generated using the collection of fit models run into the future
 - **Mean trend from last 7 days of observed cases and first week of ensemble's forecast used**
 - Outliers removed based on variances in the previous 3 weeks
 - 2 week interpolation to smooth transitions in rapidly changing trajectories
- **Outcomes:** Data driven by shift and ratio that has least error in last month of observations
 - Hospitalizations: 3 days from confirmation, 6.8% of cases hospitalized
 - Deaths: 11 days from confirmation, 1.45% of cases die



COVID-19 in Virginia: Summary

Dashboard Updated: 11/30/2022
Data entered by 5:00 PM the prior day.



Cases, Hospitalizations and Deaths					
Total Cases*		Total Hospital Admissions**		Total Deaths	
2,149,512		57,612		22,418	
(New Cases: 1,832) [^]					
Confirmed†	Probable†	Confirmed†	Probable†	Confirmed†	Probable†
1,516,580	632,932	54,043	3,569	18,591	3,827

* Includes both people with a positive test (Confirmed), and symptomatic with a known exposure to COVID-19 (Probable).
** Hospitalization of a case is captured at the time VDH performs case investigation. This underrepresents the total number of hospitalizations in Virginia.

[^]New cases represent the number of confirmed and probable cases reported to VDH in the past 24 hours.

† VDH adopted the updated CDC COVID-19 confirmed and probable surveillance case definitions on September 1st, 2021. Found here: <https://ndc.services.cdc.gov/case-definitions/coronavirus-disease-2019-2021/>

Source: Cases - Virginia Electronic Disease Surveillance System (VEDSS), data entered by 5:00 PM the prior day.

Outbreaks	
Total Outbreaks*	Outbreak Associated Cases
10,219	167,838

* At least two (2) lab confirmed cases are required to classify an outbreak.

Testing (PCR Only)	
Testing Encounters PCR Only*	Current 7-Day Positivity Rate PCR Only**
15,476,050	9.6%

* PCR refers to "Reverse transcriptase polymerase chain reaction laboratory testing."

** Lab reports may not have been received yet. Percent positivity is not calculated for days with incomplete data.

Multisystem Inflammatory Syndrome in Children	
Total Cases*	Total Deaths
180	1

*Cases defined by CDC HAN case definition: <https://emergency.cdc.gov/han/2020/han00432.asp>

Accessed 11:30am November 30, 2022

<https://www.vdh.virginia.gov/coronavirus/>



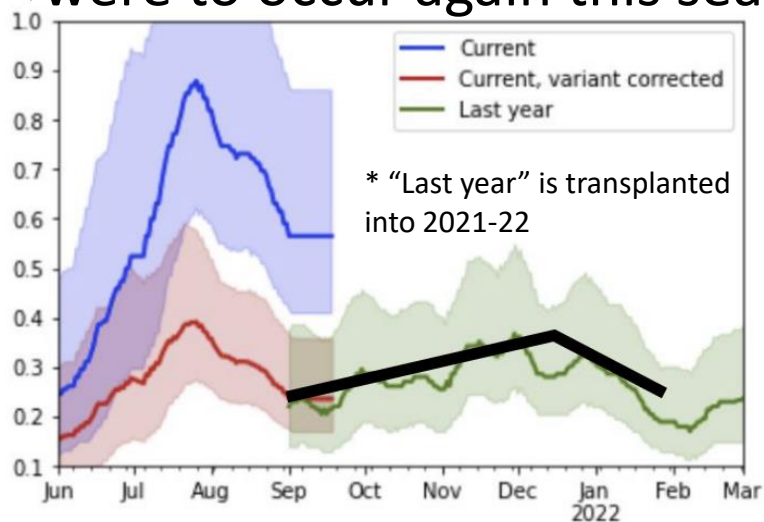
Scenarios – Transmission Conditions

- Variety of factors continue to drive transmission rates
 - Seasonal impact of weather patterns, travel and gatherings, fatigue and premature relaxation of infection control practices
- **Waning Immunity:** Omicron waning with a mean of 4 months
- **Projection Condition Ingredients:**
 - **Adaptive:** Controls remain as currently experienced into the future with NO influence from other conditions (eg seasonal, variants, etc.)
 - **Seasonal (Fall-Winter boosting):** Controls remain the same, however, seasonal forcing similar to past Fall-Winter waves is added from Sept-Feb
 - **Vaccine Booster Campaign (Booster):** Reformulated booster available this fall provides improved immunity against Omicron sub-variants
 - **New Variants (VariantX):** As of yet unidentified novel sub-variant with similar immune escape but no transmission advantage emerges 4 months after the last significant sub-variant and grows at a similar rate

Scenarios – FallWinter

September – February saw strong waves of transmission for both years

- Based on analyses of the past 2 seasons we generate a “coarse baseline transmission boost”
 - In 2021 the distribution of fitted model transmissibility was nearly identical between these periods when corrected for Delta’s increased transmissibility
- **FallWinter** captures these “transmission drivers” from the past and uses them as if they were to occur again this season

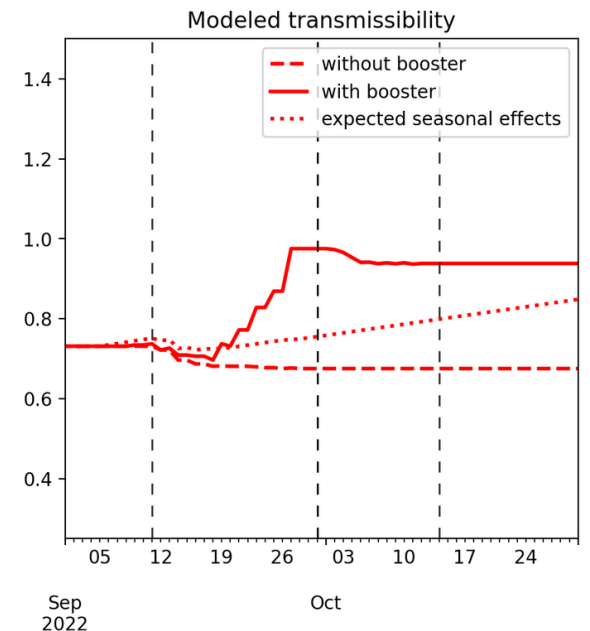


Fitting: Black line represents the coarsely fitted base transmissibility

1-Dec-22

2022 FallWinter is likely different:

With the current level of boosting the transmissibility needs to be much higher to maintain the same number of cases. The dotted line shows what transmission levels are needed to fit cases without booster and with seasonal effects.



Scenarios – Optimistic vs. Pessimistic Booster Coverage

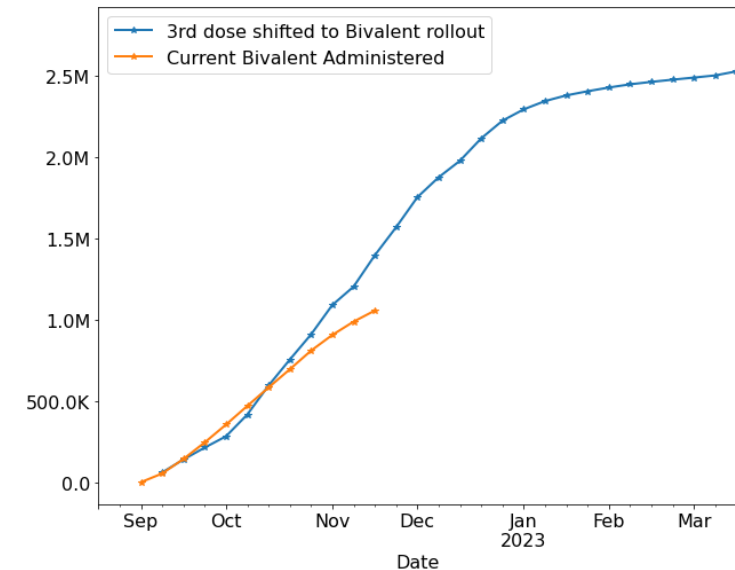
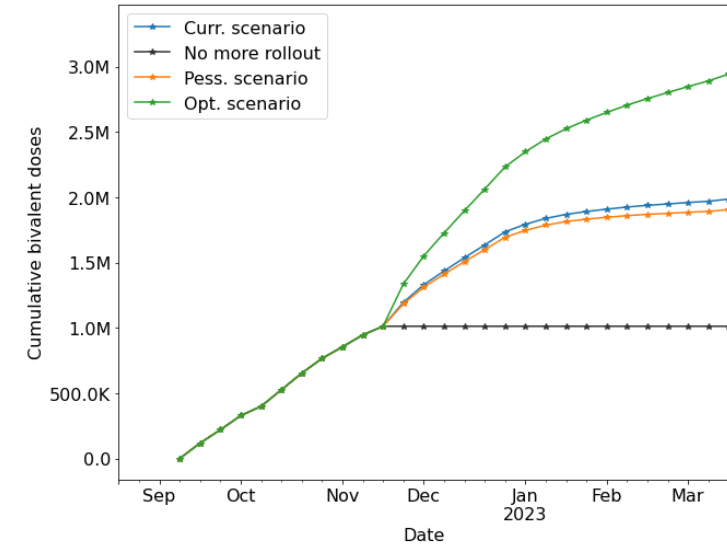
Reformulated Boosters available now

- Assume vax efficacy for BA.4/5 and previous variants is 80% against symptomatic illness
- Campaign follows current ground truth to present
- Variant X has same immune escape to these vaccines as against BA.5 (33%)

Current pace: Follows 3rd dose rollout, but maintains current pace relative to it (eg if slower, same slower rate continues into future)

Optimistic pace: 25% higher than previous 3rd dose schedule

No More: No further Bivalent boosters administered

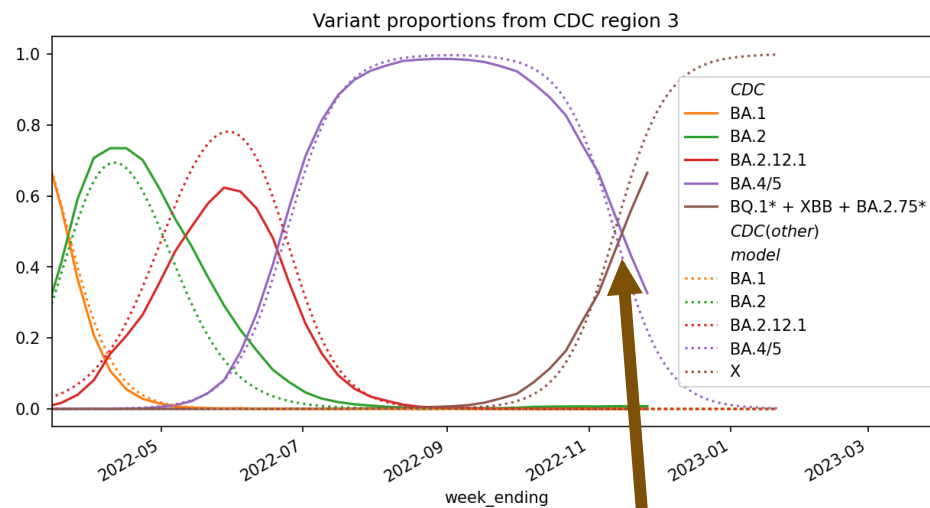


Scenarios – Variant X

Omicron sub-variants seem to be emerging and then dominating with some regularity

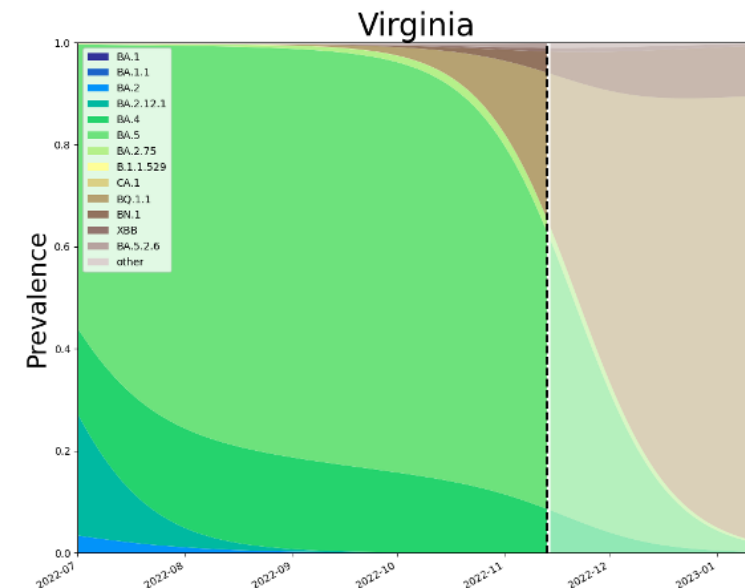
- An increasingly complex soup of variants with demonstrated growth advantages in other countries and states continues to grow
- BQ.1.1, XBB, and others have shown evidence of significant immune escape, BQ.1.1 currently growing quickly in US and VA, it remains possible that several may simultaneously expand
- **VariantX** represents the next variant or the potential swarm of several. We assume similar growth and level of immune escape against previous sub-variants as BA.4/5 (same transmissibility and 30% immune escape against BA.4/5, higher for other sub-variants).

Sub-Variants with Fitted Prevalences and Hypothetical Future waves



1-Dec-22

Variant X reached 50% on Nov 12th



Projection Scenarios – Combined Conditions

Name	Txm	Variant	Booster	Description
Adaptive	C	SQ	Current	Likely trajectory based on conditions remaining similar to the current experience, includes immune escape due to Omicron
Adaptive-FallWinter	FallWinter	SQ	Current	Like Adaptive, with seasonal forcing of FallWinter added on
Adaptive-VariantX	C	X	Current	Like Adaptive, with emergence of a speculative unknown variant 4 months after BA.4/5 with similar level of immune escape and equal transmissibility
Adaptive-VariantX-FallWinter	FallWinter	X	Current	Like Adaptive-VariantX but with the seasonal force of FallWinter added on
Adaptive-VariantX-FallWinter-OptBooster	FallWinter	X	Optimistic	Like Adaptive-VariantX-Fall Winter but with Optimistic Booster (25% more than 3 rd dose rollout)
Adaptive-VariantX-FallWinter-NoMoreBooster	FallWinter	X	No More	Like Adaptive-VariantX-FallWinter but with no additional Booster doses

Transmission:

C = Current levels persist into the future

FallWinter = Transmission rates learned from Sept through February of past seasons are estimated and added as a seasonal boosting to baseline transmission rates

Variant:

SQ = Status quo prevalences remain the same (e.g. no significant major driving of transmission anticipated)

X = Novel sub-variant scenario, new variant emerges reaches dominance in near term, 30% immune escape

Booster:

Current = Current pace relative to 3rd dose rollout is maintained in the future

Optimistic = Starting this week, additional 25% over the 3rd dose rollout is maintained into the future

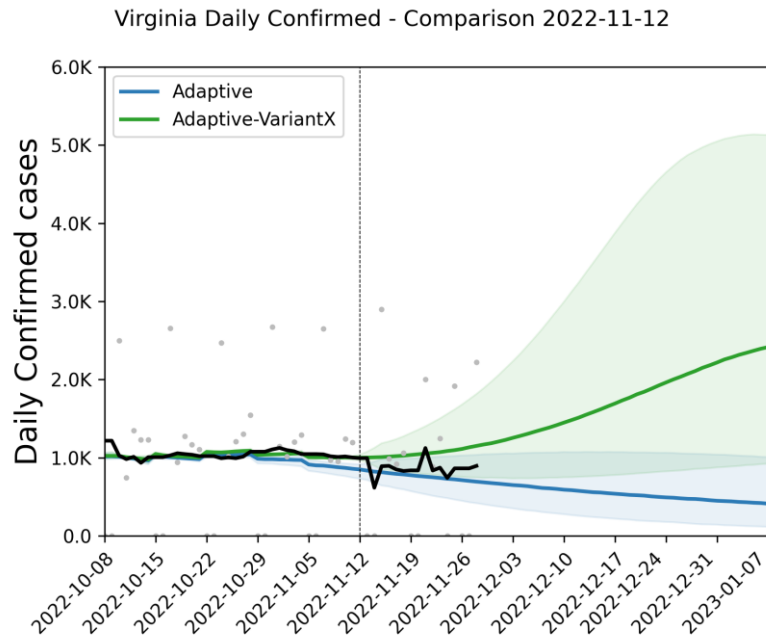
No More = Starting this week, no additional doses of the booster are administered

Model Results

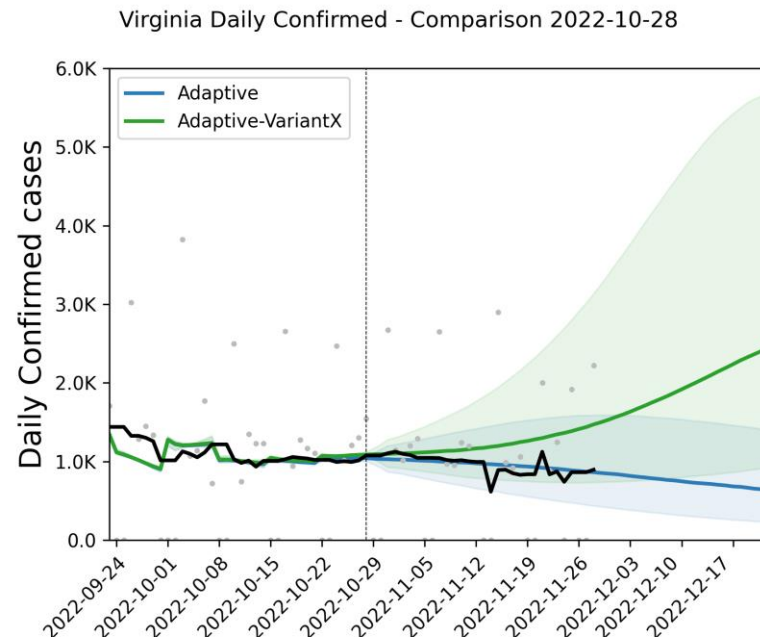
Previous projections comparison - Cases

- Previous projections continue to track observed cases
- Projection from 2 weeks ago projected plateau a week after cases started to plateau
- Projection from 4 weeks ago projected slower decline better capturing recent slowing
- Projection from early July remains eerily on track

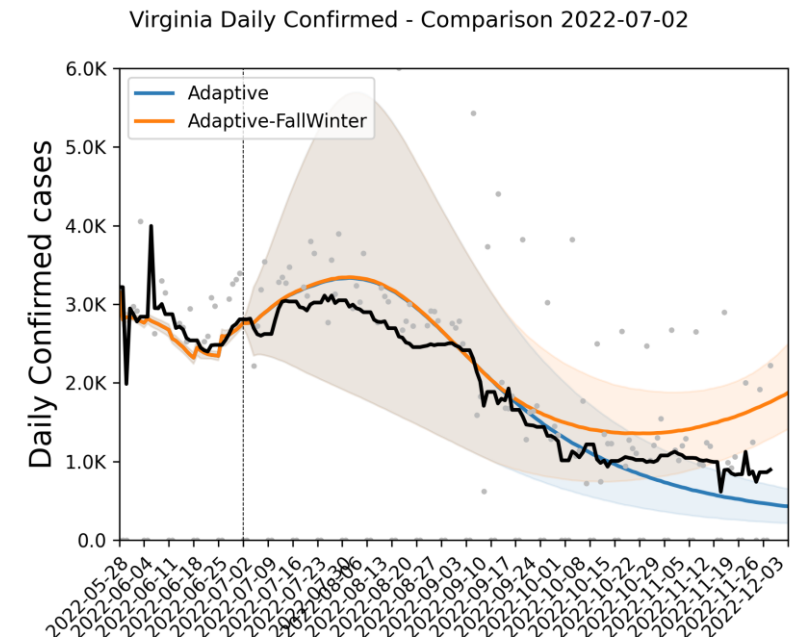
Previous round (2 weeks ago)



Projection from 4 weeks ago



Projection from 3 months ago

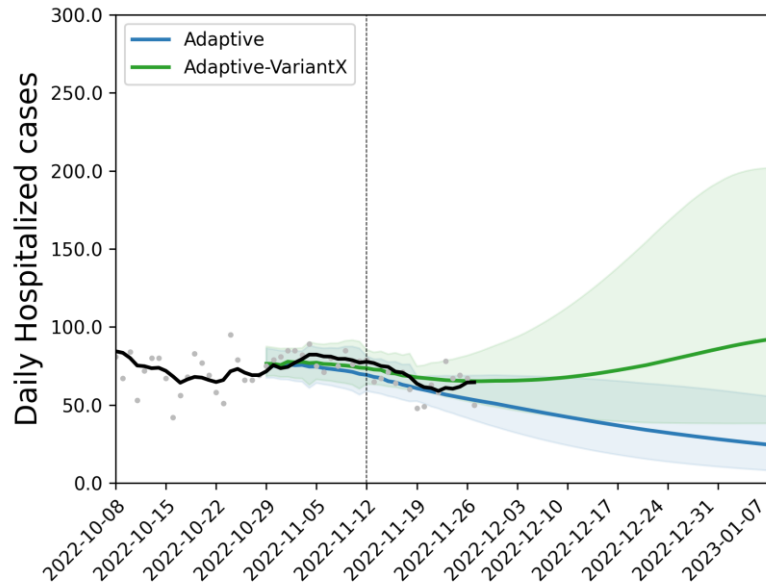


Previous projections comparison - Hospitalizations

- Previous projections have tracked observed hospitalizations well
- Projection from 2 weeks ago projected continued decline, missed bump
- Projection from early July anticipated a plateau has tracked reasonably well up to present

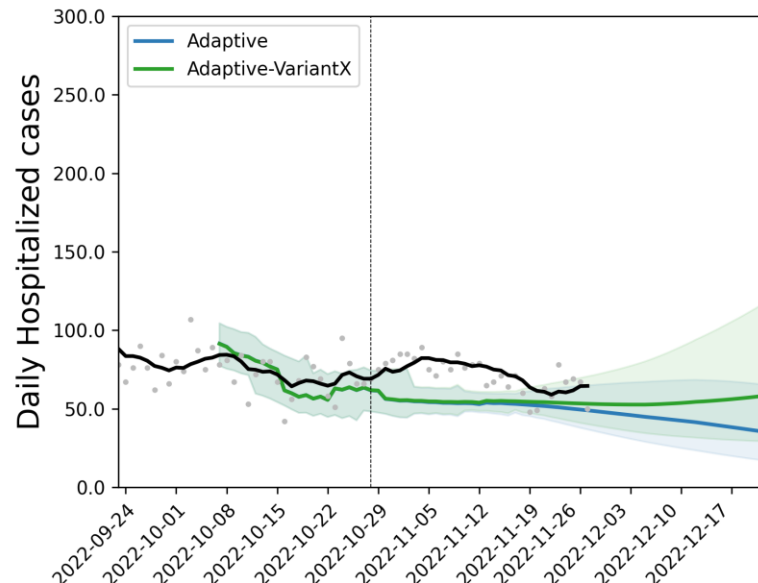
Previous round (2 weeks ago)

Virginia Daily Hospitalized - Comparison 2022-11-12



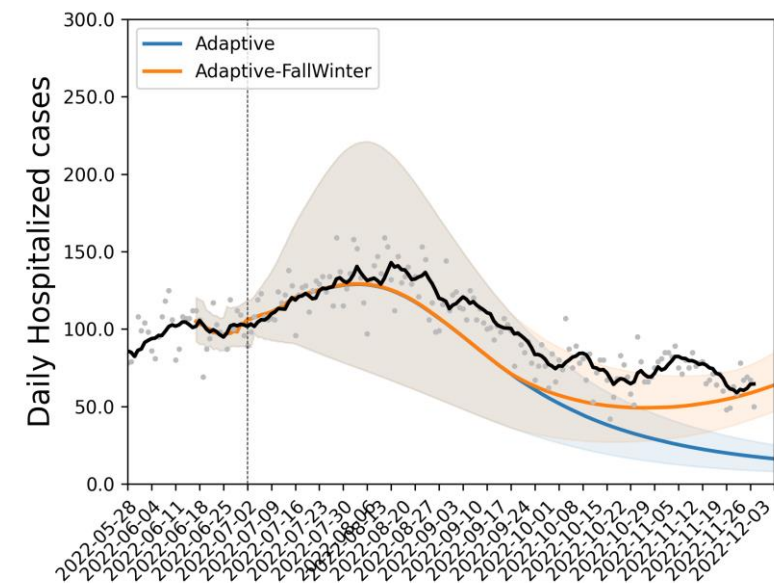
Projection from 4 weeks ago

Virginia Daily Hospitalized - Comparison 2022-10-28



Projection from 3 months ago

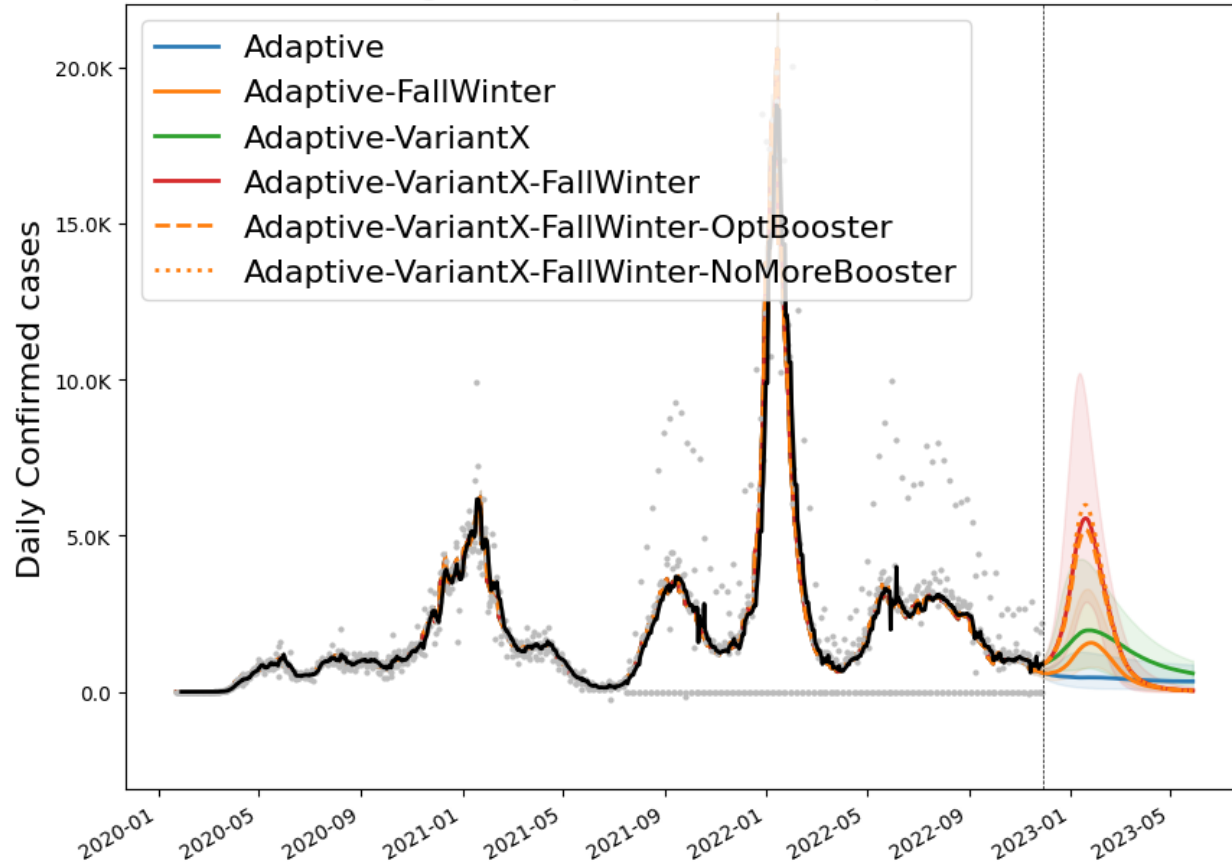
Virginia Daily Hospitalized - Comparison 2022-07-02



Outcome Projections

Confirmed cases

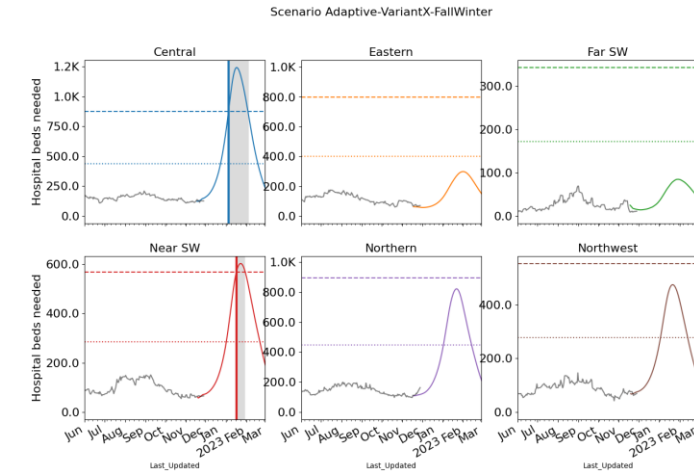
Virginia Daily Confirmed - Comparison



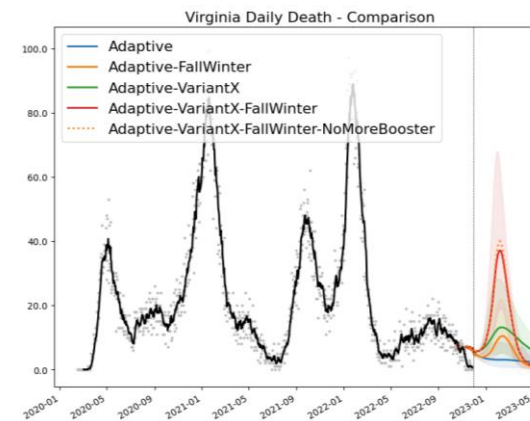
* without surveillance correction VariantBA2 peaked over 10K in July



Estimated Hospital Occupancy

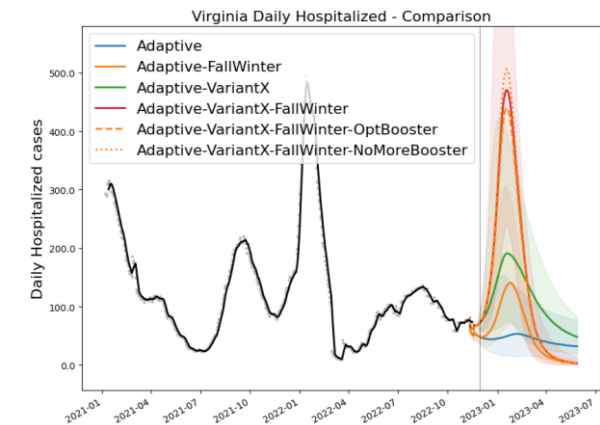


Daily Deaths



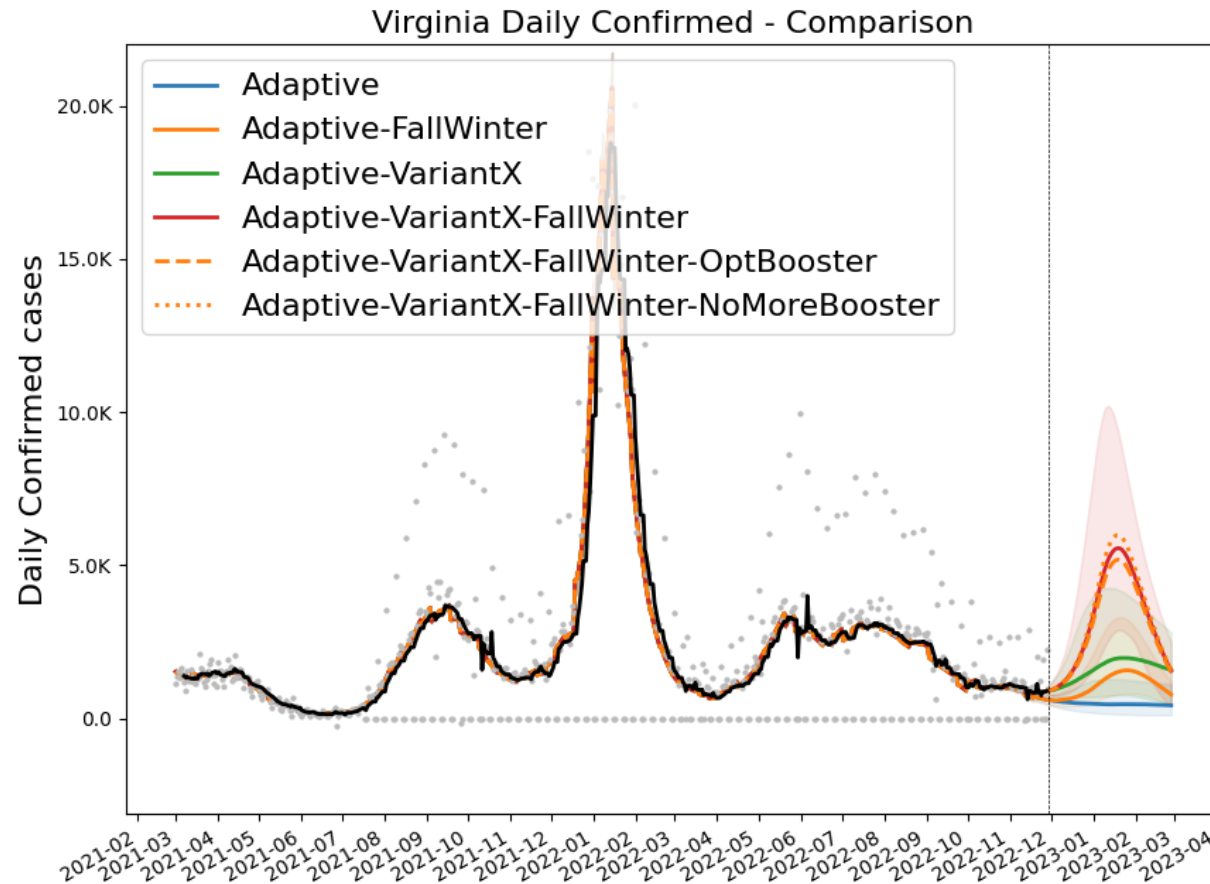
Death ground truth from VDH "Event Date" data, most recent dates are not complete

Daily Hospitalized



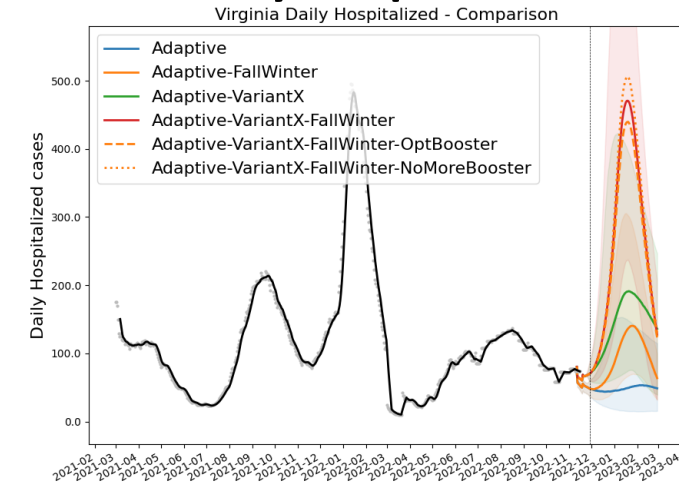
Outcome Projections – Closer Look

Confirmed cases

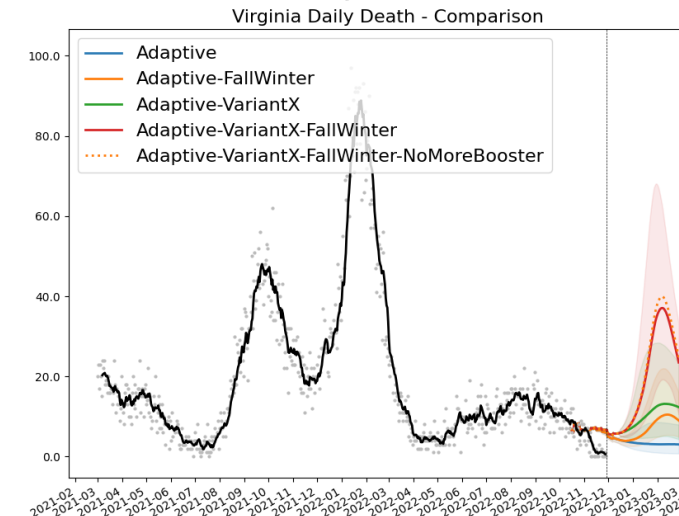


* without surveillance correction VariantBA2 peaked over 10K in July

Daily Hospitalized



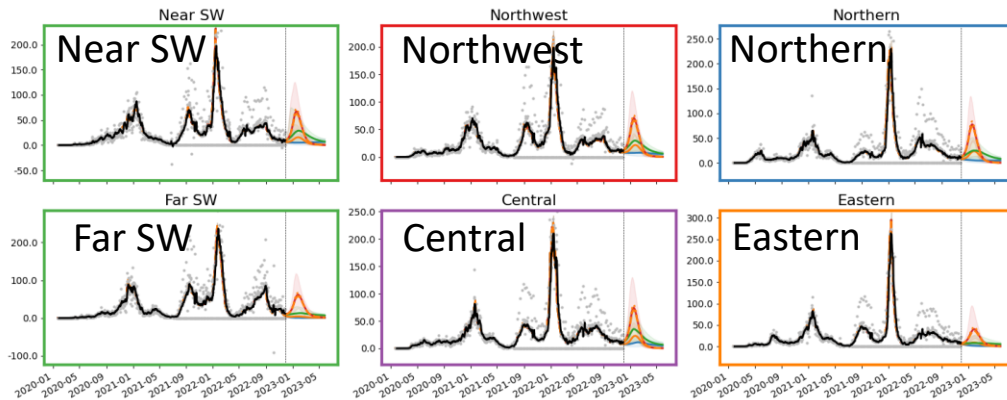
Daily Deaths



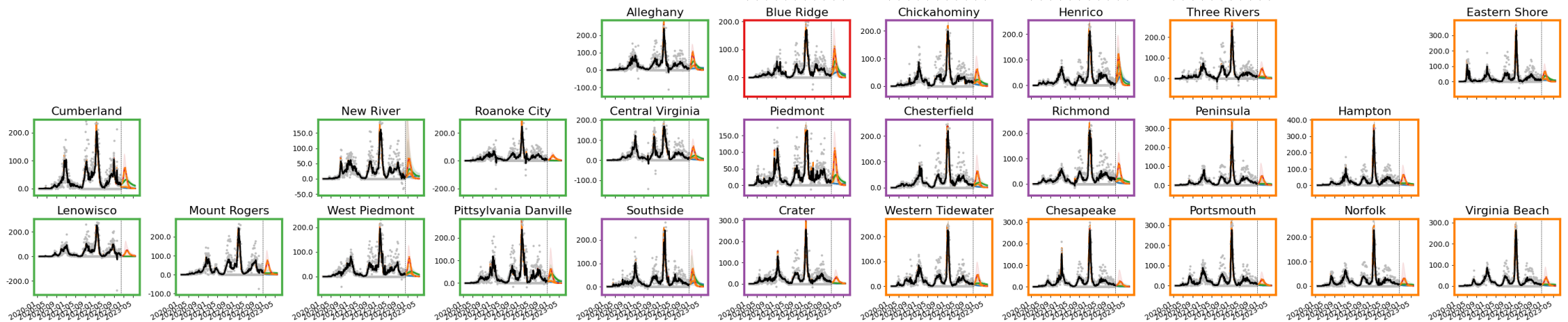
Death ground truth from VDH "Event Date" data, most recent dates are not complete

Detailed Projections: Cases for All Scenarios

Projections by Region



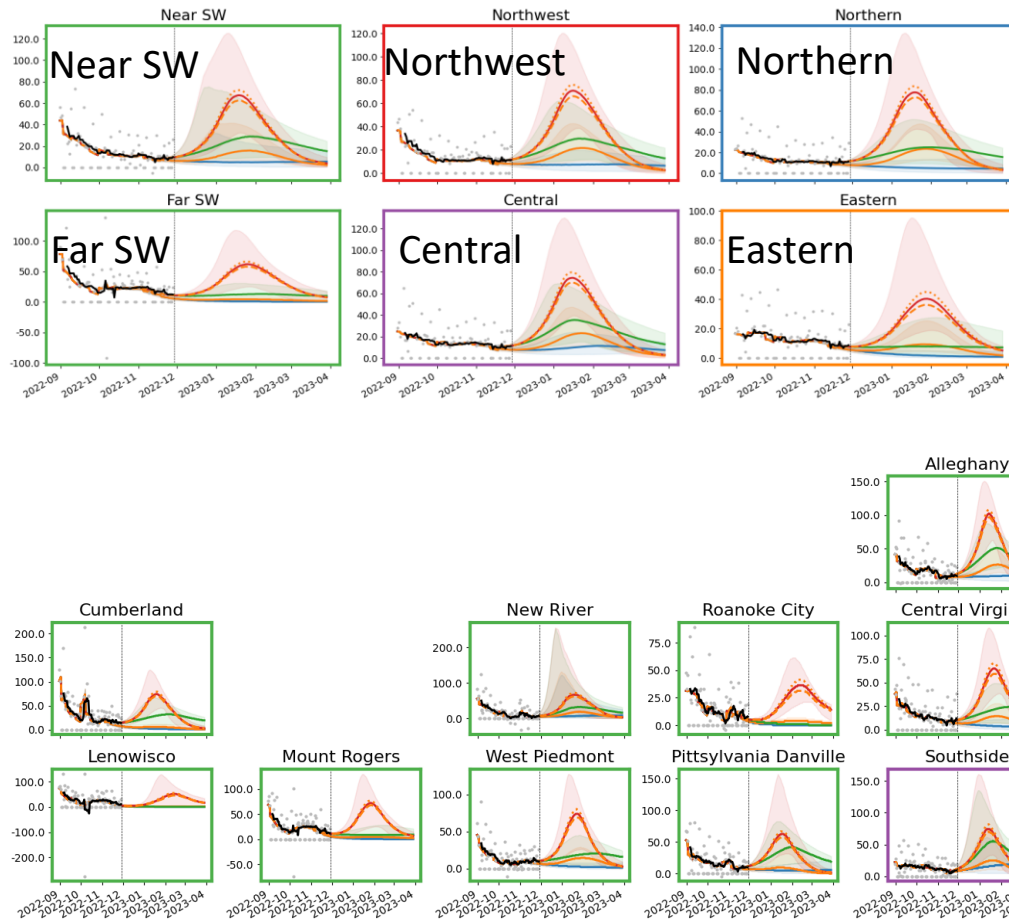
Projections by District



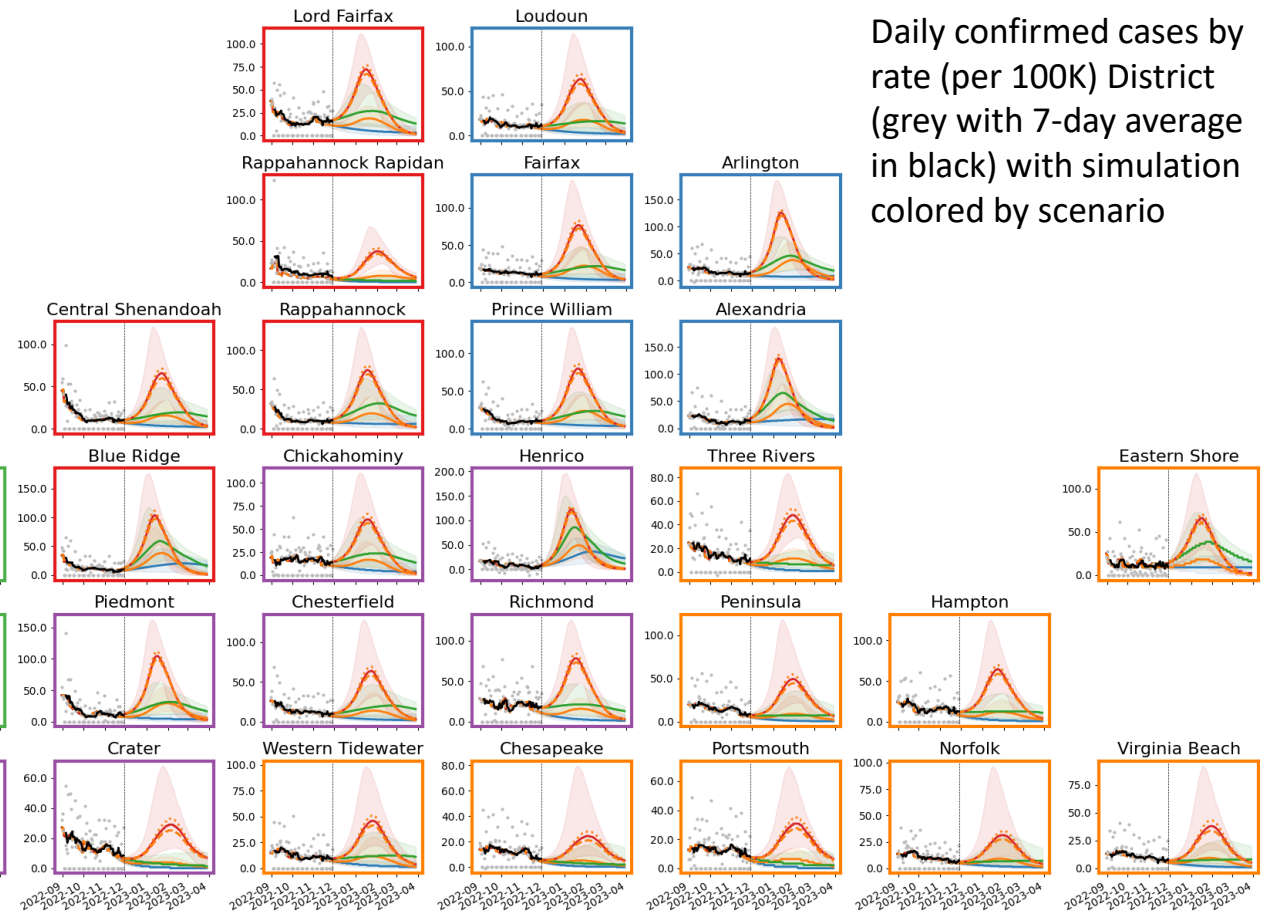
Daily confirmed cases)
by rate (per 100K)
District (grey with 7-day
average in black) with
simulation colored by
scenario

Detailed Projections: Cases for All Scenarios - Closer Look

Projections by Region



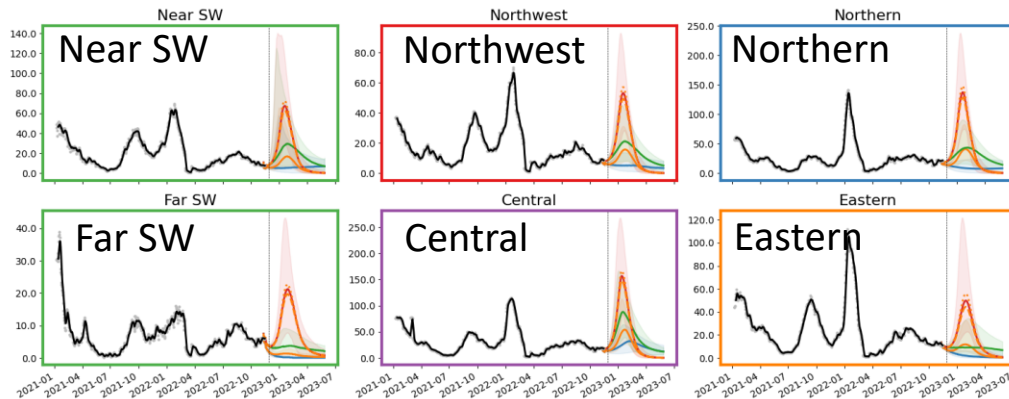
Projections by District



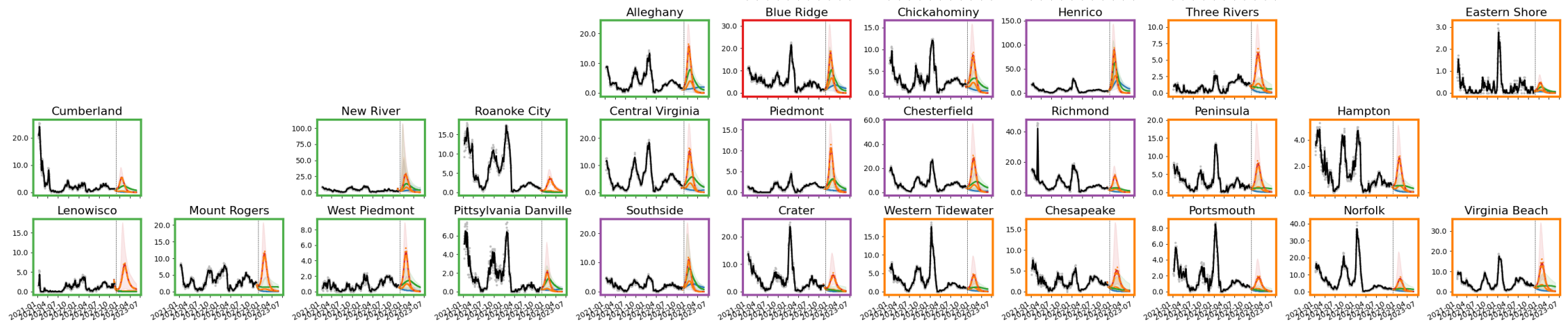
Daily confirmed cases by rate (per 100K) District (grey with 7-day average in black) with simulation colored by scenario

Detailed Projections: Hospitalizations for All Scenarios

Projections by Region



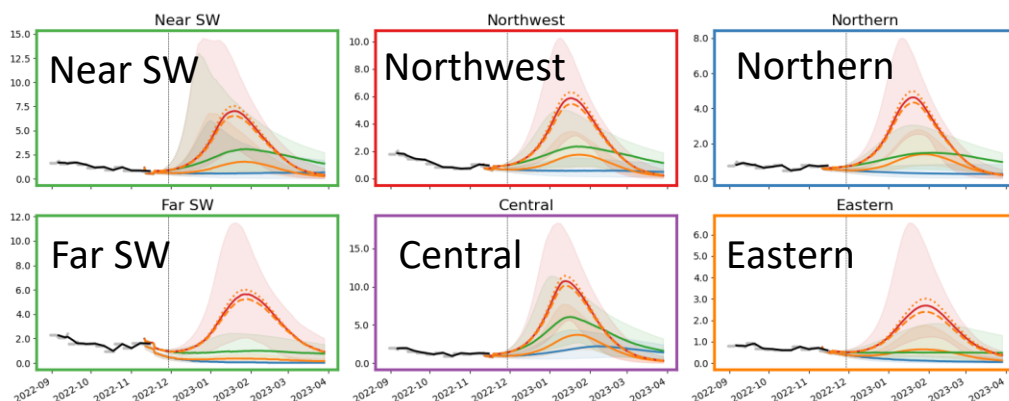
Projections by District



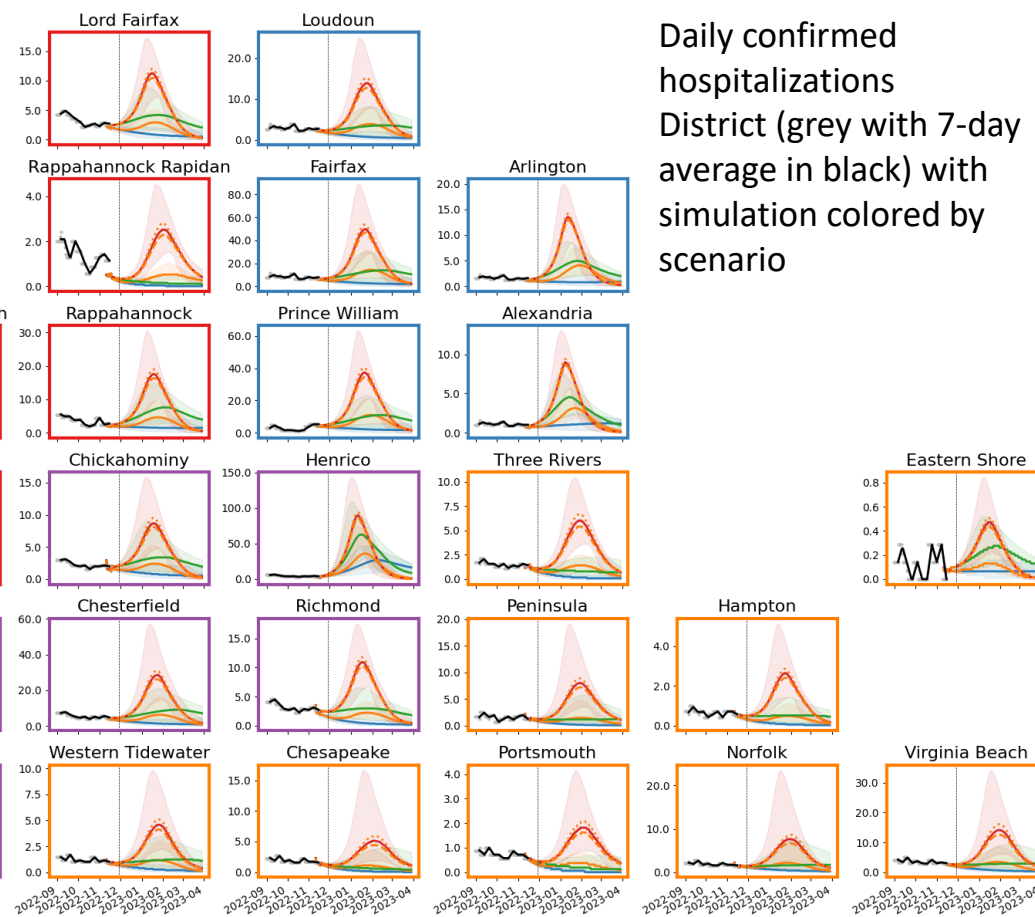
Daily confirmed hospitalizations District (grey with 7-day average in black) with simulation colored by scenario

Detailed Projections: Hosps for All Scenarios - Closer Look

Projections by Region



Projections by District

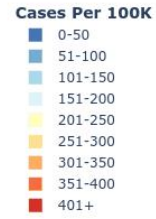
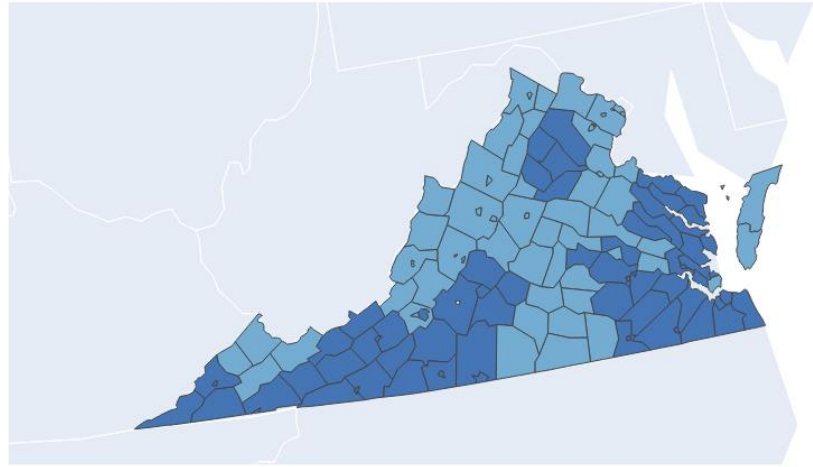


Daily confirmed hospitalizations
District (grey with 7-day average in black) with simulation colored by scenario

Adaptive

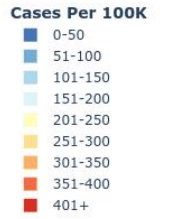
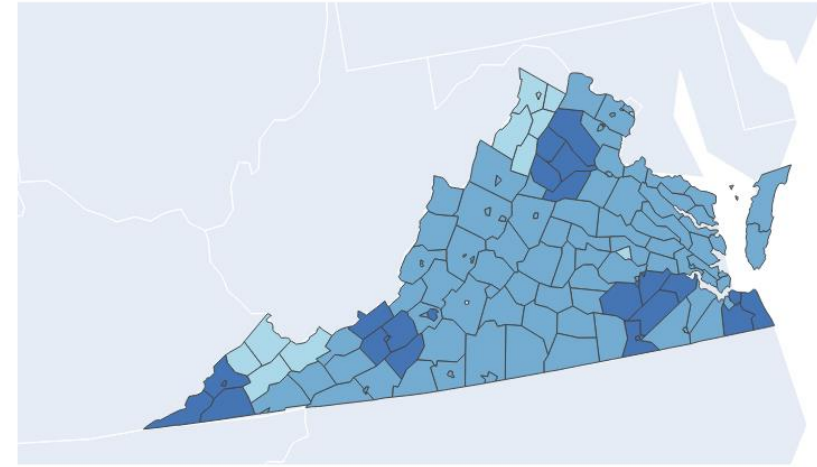
Adaptive

Weekly Projections (Adaptive) 23-Nov-2022



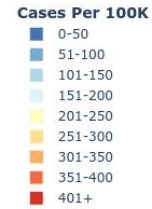
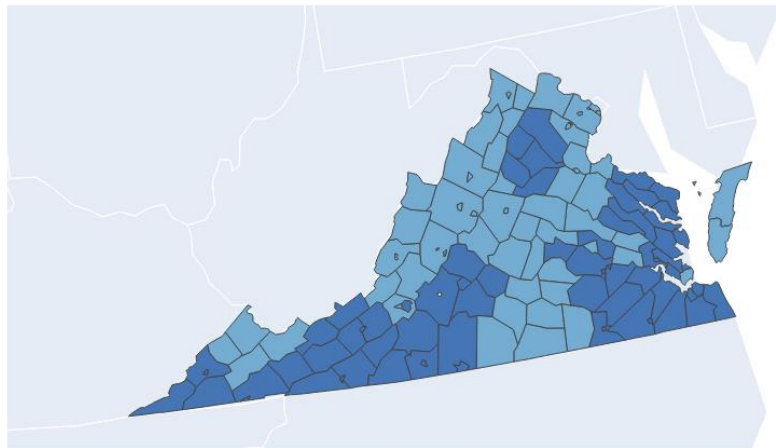
VariantX

Weekly Projections (Adaptive-VariantX) 23-Nov-2022

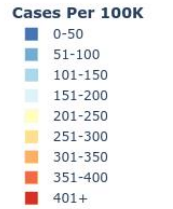
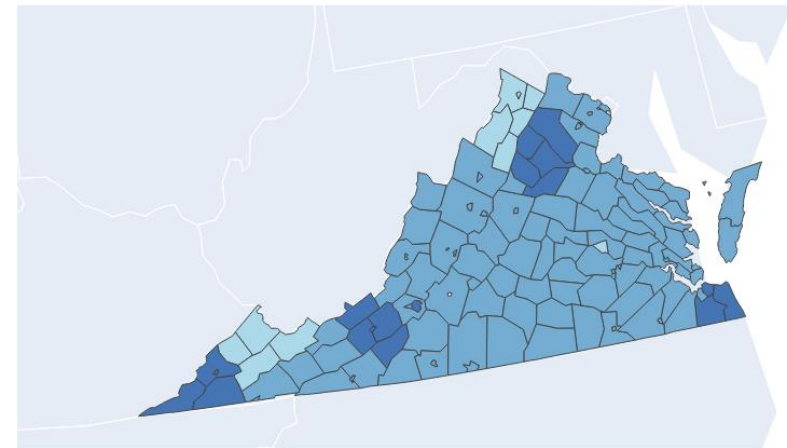


Adaptive-Fall-Winter

Weekly Projections (Adaptive-FallWinter) 23-Nov-2022



Weekly Projections (Adaptive-VariantX-FallWinter) 23-Nov-2022

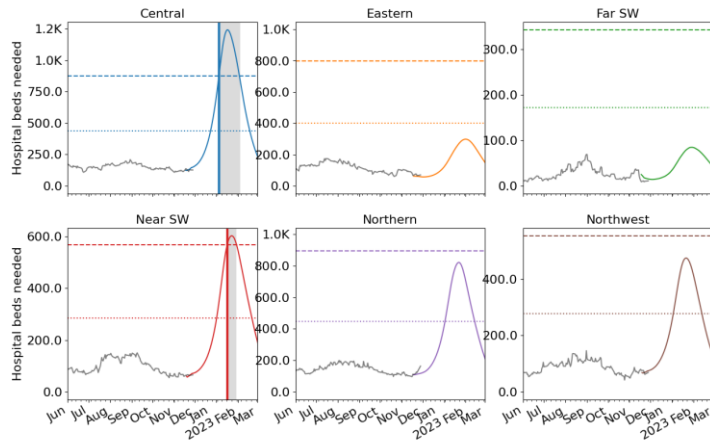


Hospital Demand and Bed Capacity by Region

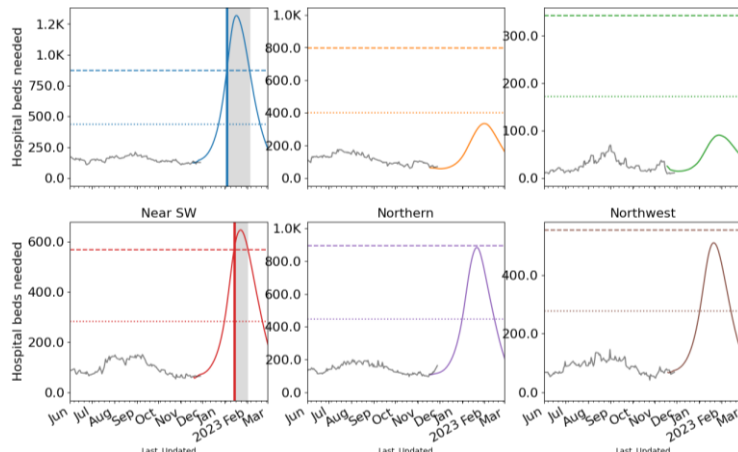
Capacities by Region

COVID-19 capacity ranges from 80% (dots) to 120% (dash) of total beds

Adaptive – VariantX & Fall Winter



Adaptive – VariantX & Fall Winter No More Booster



Last Updated
1-Dec-22

Length of Stay more variable with Omicron, occupancy projections may vary as a result, ad-hoc estimation performed per region

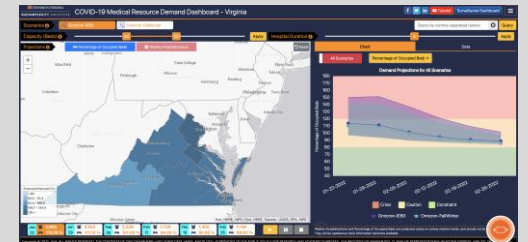
Estimated LOS shortened slightly to better fit observed data

Projections show continued declines and with expanded capacities and adjusted length of stay, no capacities exceeded

Length of Stay Estimates

Central	6
Eastern	6
Far SW	4
Near SW	9
Northern	5
Northwestern	9

Interactive Dashboard
with regional
projections



<https://nssac.bii.virginia.edu/covid-19/vmrddash/>

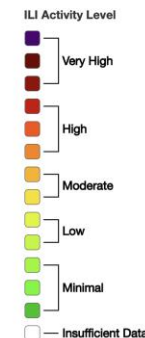
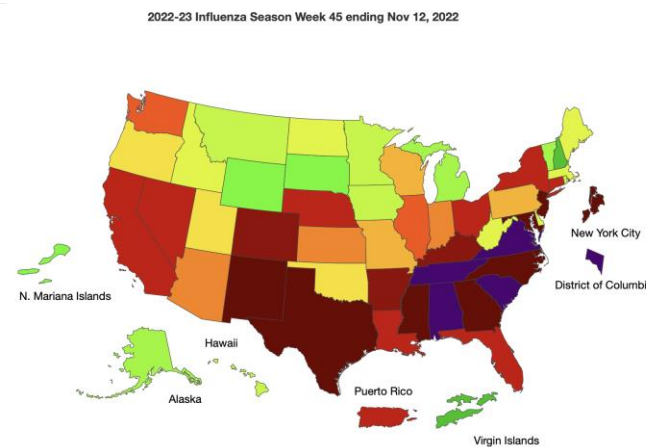
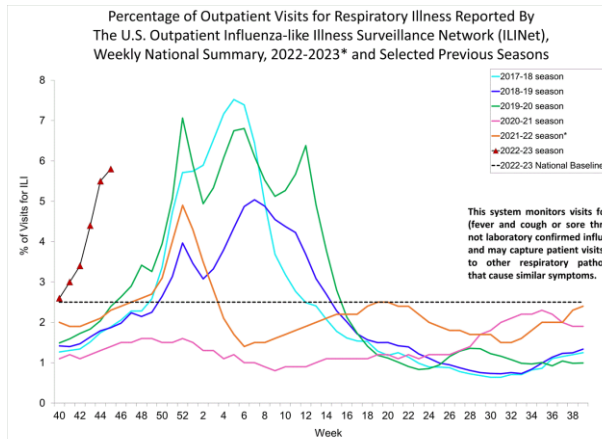
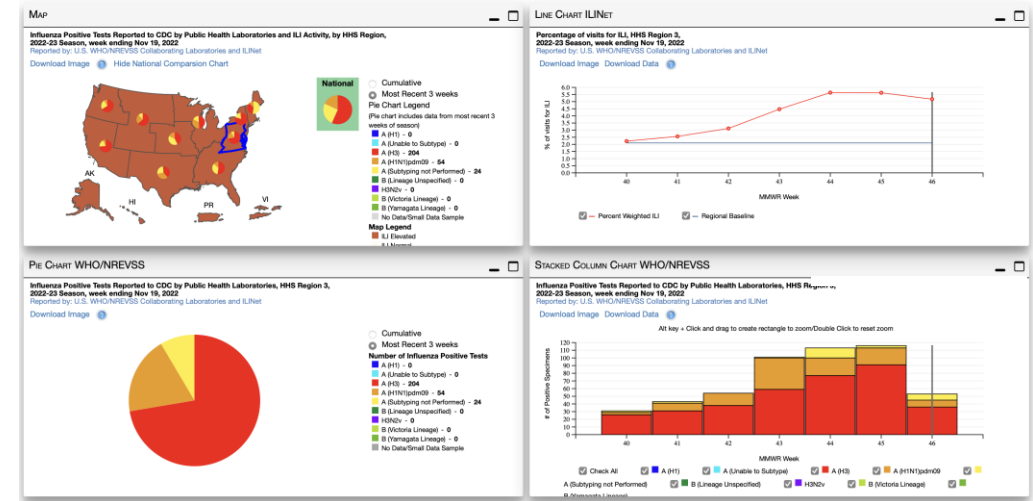
Influenza Update

Current Influenza Situation – ILI Activity

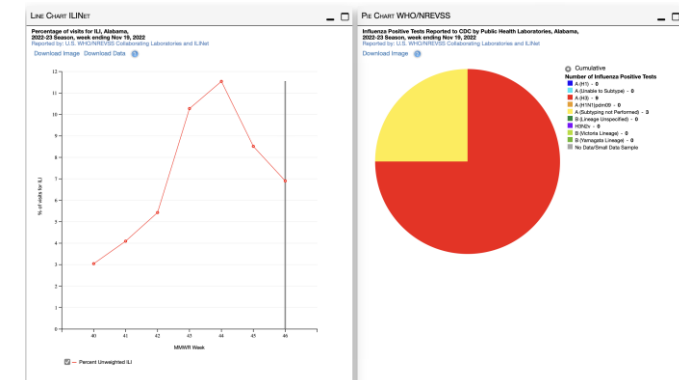
Influenza Activity is Higher than Usual

- Virginia at “Very High” activity along with much of the Southeast, with “High” activity shifting out across US
- In VA ILI Activity has receded from a high near ~12% of visits for ILI back to 7% this past week
- National ILI activity at highest before December since 2009 pandemic year which peaked in mid-Oct
- After starting with high proportions of H3N2 typed influenza, H1N1pdm09 now represents ~1/5 of all infections nationally

Region 3



Virginia

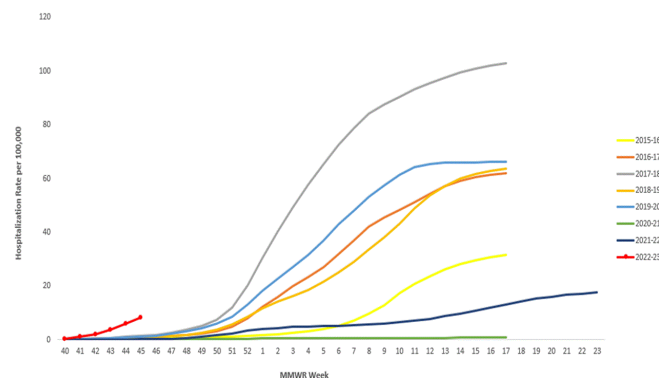


Current Influenza Situation - Hospitalizations

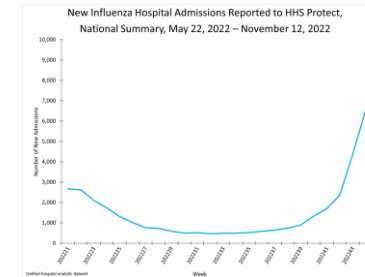
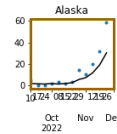
Influenza A hospitalizations continue rapid growth

- National level of influenza hospitalizations
- Nearly all states have doubled their hospitalizations due to influenza in the last couple weeks
- Virginia shows leveling off in the last weeks

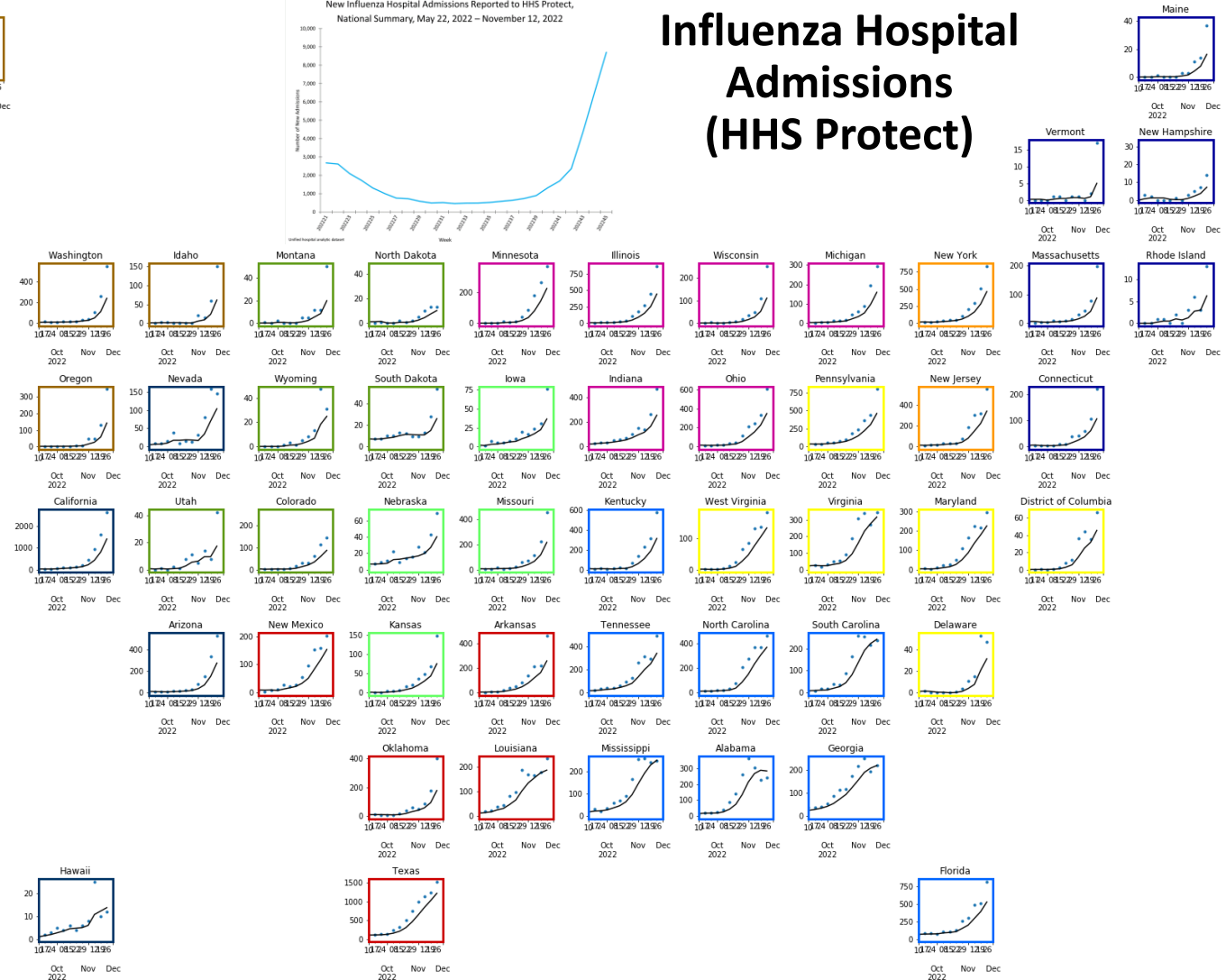
Cumulative Rate of Laboratory-Confirmed Influenza Hospitalizations among cases of all ages, 2015-16 to 2022-23, MMWR Week 45



**In this figure, cumulative rates for all seasons prior to the 2022-23 season reflect end-of-season rates. For the 2022-23 season, rates for recent hospital admissions are subject to reporting delays. As hospitalization data are received each week, prior case counts and rates are updated accordingly.



Influenza Hospital Admissions (HHS Protect)

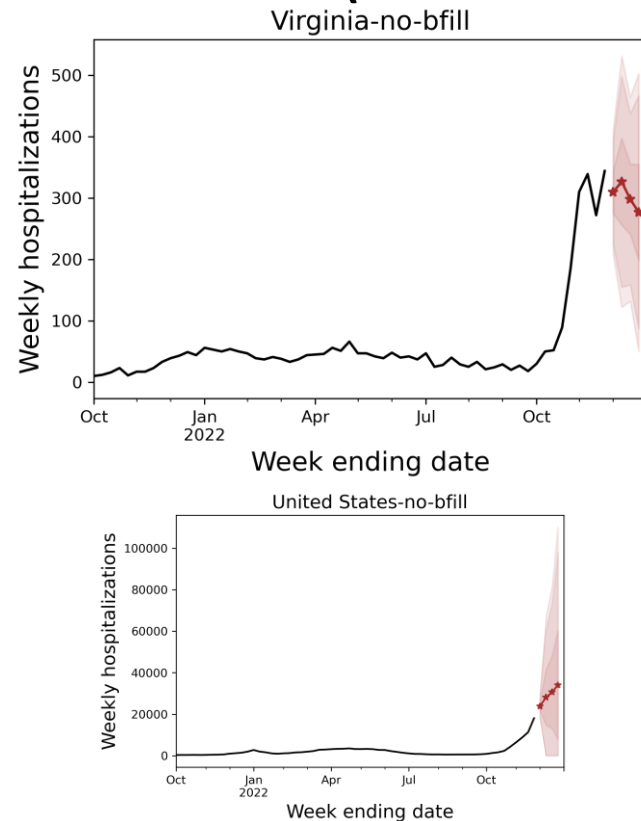


Current Influenza Hospitalization Forecast

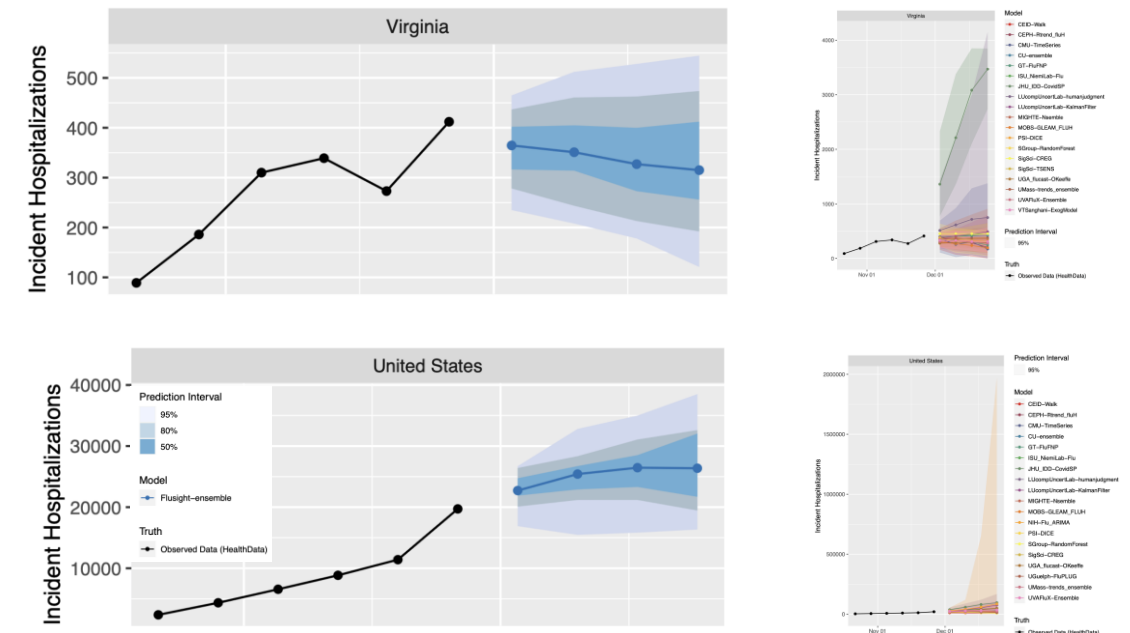
Statistical models for submitting to CDC FluSight forecasting challenge

- Similar to COVID-19 case forecasts, uses a variety of statistical and ML approaches to forecast weekly hospital admissions for the next 4 weeks for all states in the US

Hospital Admissions for Influenza and Forecast for next 4 weeks (UVA ensemble)



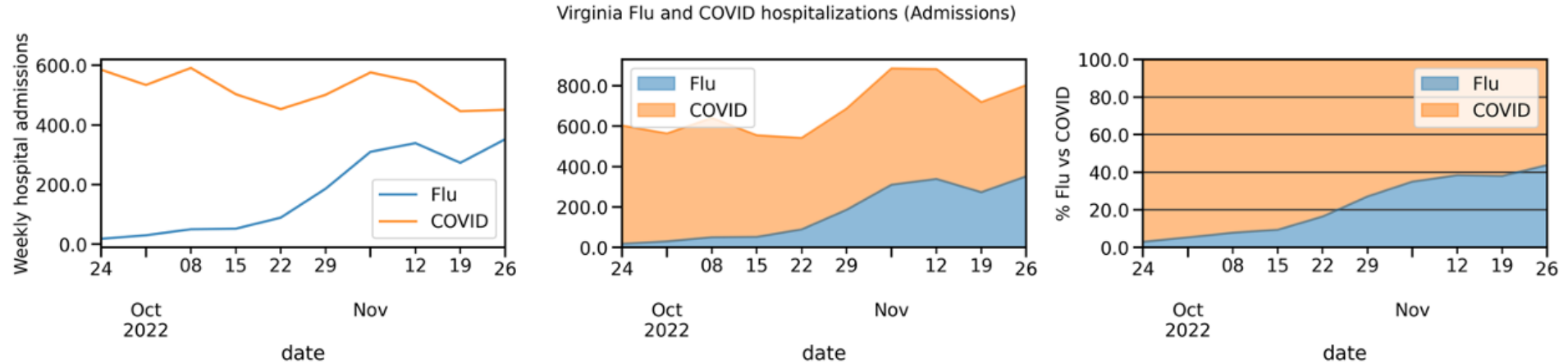
Hospital Admissions for Influenza and Forecast for next 4 weeks (CDC FluSight Ensemble)



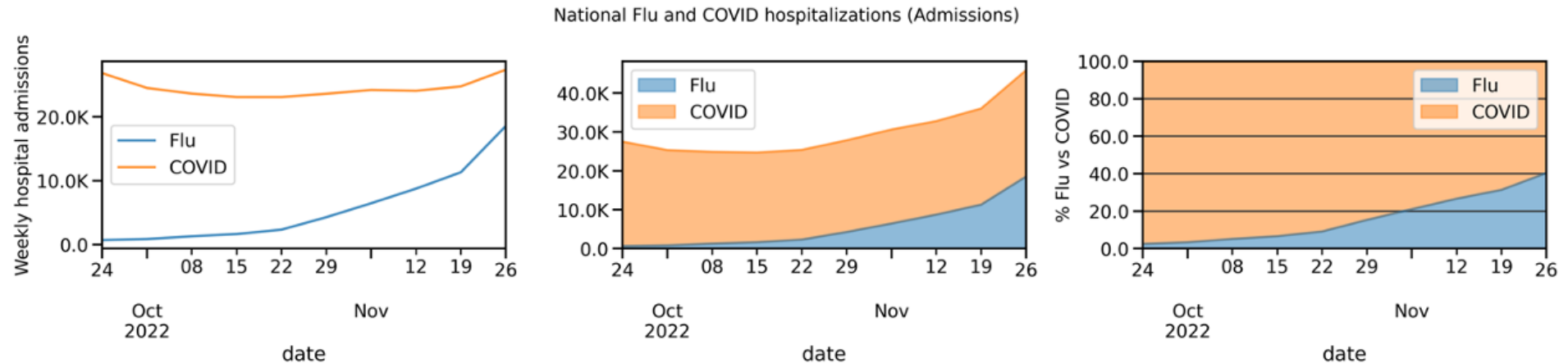
Current Combined Hospitalizations (COVID-19 & Influenza)

COVID-19 and Influenza Weekly Hospitalizations (HHS Protect)

Virginia



USA



Combined ILI and COVID-19 Hospitalizations

Ensemble methodology that combines the Adaptive with machine learning and statistical models such as:

- Autoregressive (AR, ARIMA) , Neural networks (LSTM), Kalman filtering (EnKF), G-model (phase), Holt-Winters

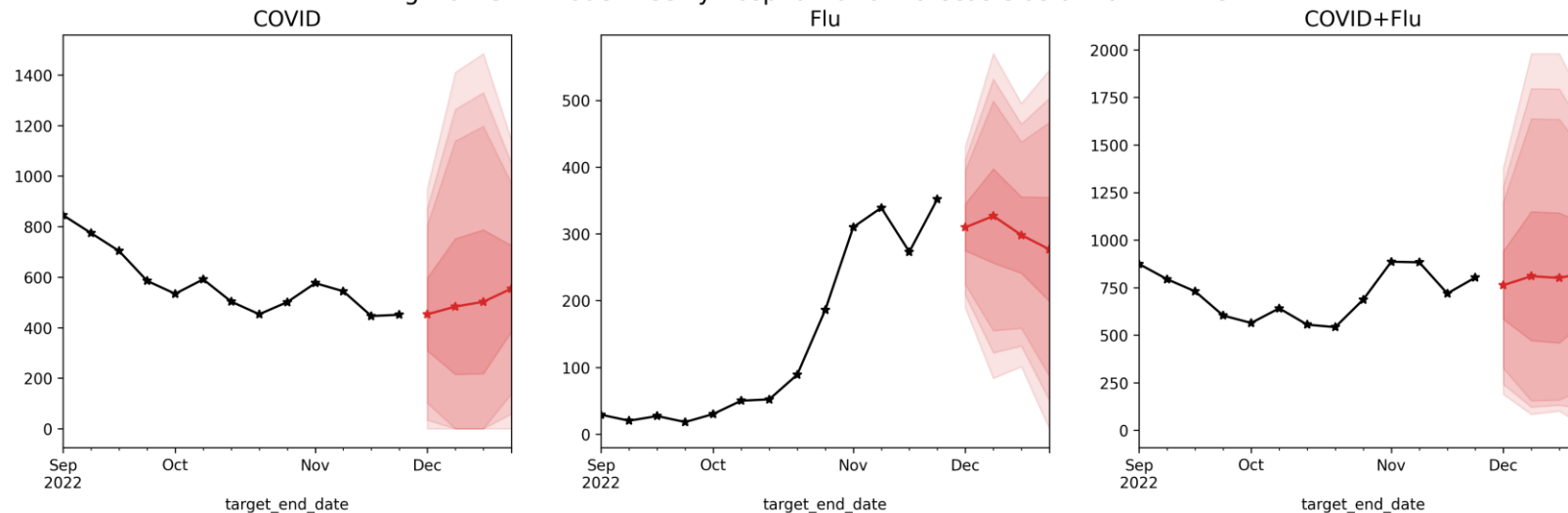
Weekly forecasts of hospitalizations done at state level.

Models chosen because of their track record in disease forecasting and to increase diversity and robustness.

Both are regularly submitted to CDC Forecast Hubs

Weekly Hospitalizations Short-term COVID-19 and Influenza Forecasts

Virginia - UVA model weekly hospitalization forecasts as of 2022-11-28



National Modeling Hub Updates

Scenario Modeling Hub – COVID-19 (Round 16)

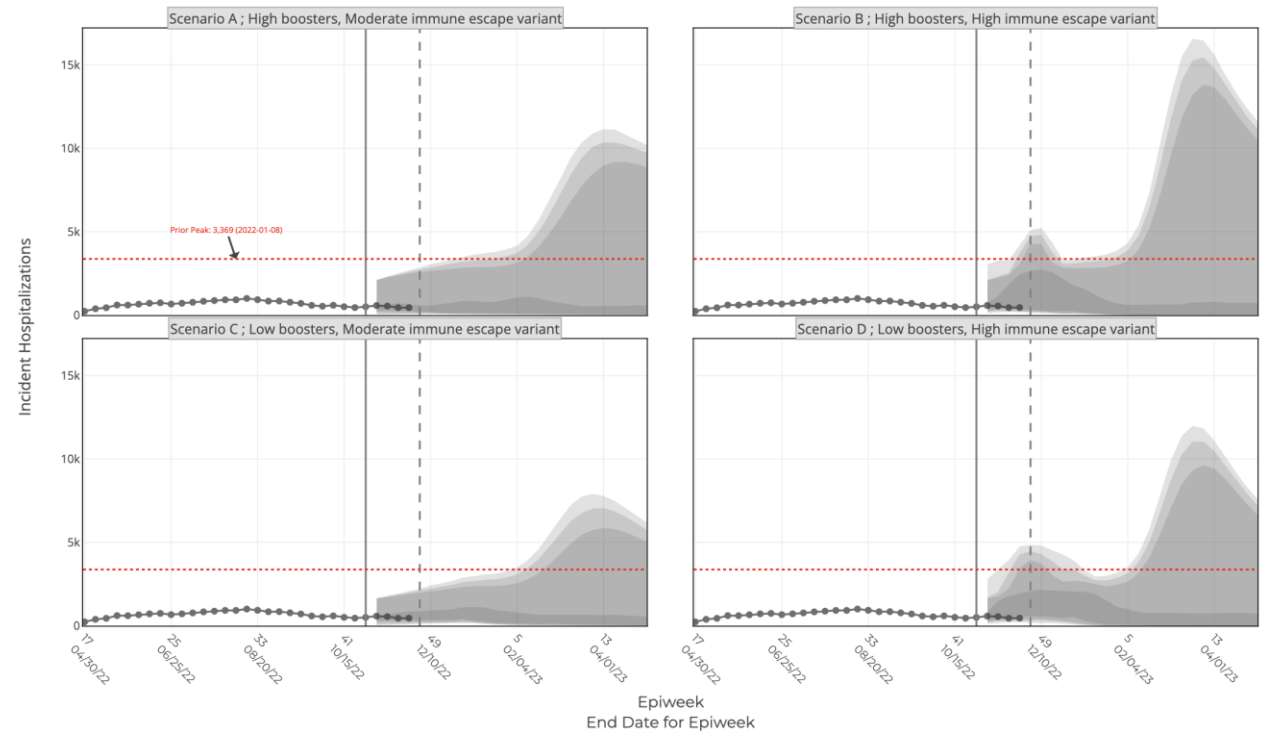
Collaboration of multiple academic teams to provide national and state-by-state level projections for 4 aligned scenarios

- Round 16 results published

	"Level 5" Variants	"Level 6/7" Variants
Accelerating uptake levels of reformulated boosters	Scenario A "Level 5" Variants - Variants have a 25% immune escape from BA.5.2 - Seeding based on combined observed prevalence of Level 5 variants at the start of the projection period - No change in severity given symptomatic infection Accelerating uptake levels of reformulated boosters, with coverage plateauing at 90% of flu vaccination levels by February 1st, 2023 - Teams are free to use available data and information from current and previous rollouts as they see fit to define rates - Teams should assume increasing uptake through October and November as necessary to reach the projected February 1st, 2022 plateau	Scenario B "Level 6/7" Variants - Variants have a 50% immune escape from BA.5.2 - Seeding based on combined observed prevalence of Level 6 and 7 variants at the start of the projection period - No change in severity given symptomatic infection Accelerating uptake levels of reformulated boosters, with coverage plateauing at 90% of flu vaccination levels by February 1st, 2023 - Teams are free to use available data and information from current and previous rollouts as they see fit to define rates - Teams should assume increasing uptake through October and November as necessary to reach the projected February 1st, 2022 plateau
Current uptake levels of reformulated boosters	Scenario C "Level 5" Variants - Variants have a 25% immune escape from BA.5.2 - Seeding based on combined observed prevalence of Level 5 variants at the start of the projection period - No change in severity given symptomatic infection Current uptake levels of reformulated boosters, with coverage plateauing at booster 1 levels by the end of the simulation - Teams are free to use available data and information from current and previous rollouts as they see fit to define rates - Based on current rates, plateau date is flexible as long as it occurs before the end of the simulation (Teams can adjust rates up if needed to achieve adequate coverage by target date)	Scenario D "Level 6/7" Variants - Variants have a 50% immune escape from BA.5.2 - Seeding based on combined observed prevalence of Level 6 and 7 variants at the start of the projection period - No change in severity given symptomatic infection Current uptake levels of reformulated boosters, with coverage plateauing at booster 1 levels by the end of the simulation - Teams are free to use available data and information from current and previous rollouts as they see fit to define rates - Based on current rates, plateau date is flexible as long as it occurs before the end of the simulation (Teams can adjust rates up if needed to achieve adequate coverage by target date)

<https://covid19scenariomodelinghub.org/viz.html>

Projected Incident Hospitalizations by Epidemiological Week and by Scenario for Round 16 - Virginia
(- Projection Epiweek; -- Current Week)



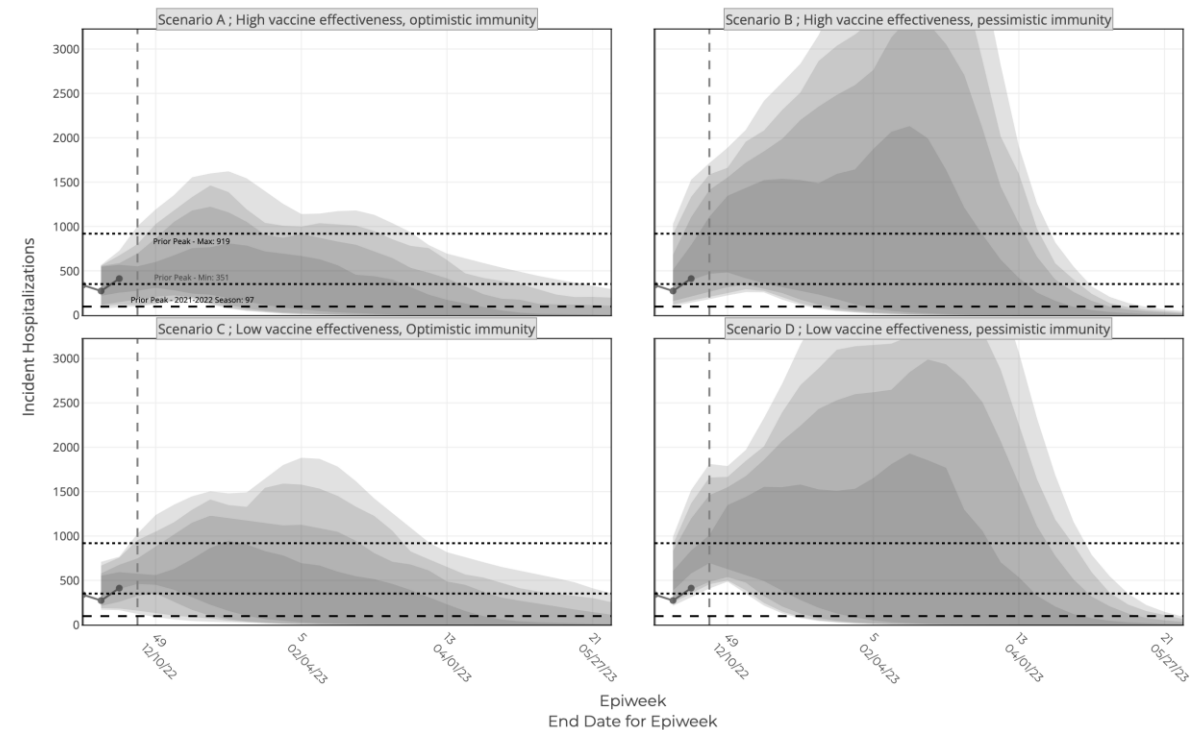
Scenario Modeling Hub – Influenza (Round 2 - prelim)

Collaboration of multiple academic teams to provide national and state-by-state level projections for 4 aligned scenarios

- Round 2 is more calibrated and similar to Round 1 but not complete

<https://fluscenariomodelinghub.org/viz.html>

Projected Incident Hospitalizations by Epidemiological Week and by Scenario for Round 2 - Virginia
(- Projection Epiweek; -- Current Week)



	Optimistic flu prior immunity	Pessimistic flu prior immunity
High Vaccine Effectiveness	Scenario A Optimistic flu prior immunity - No impact of missed flu seasons due to the COVID-19 pandemic on prior immunity.* - Same amount of prior immunity as in a typical, pre-COVID19 pandemic prior season. High Vaccine Effectiveness - VE = 50% against medically attended influenza illnesses and hospitalizations (comparable to 2015-16 season).	Scenario B Pessimistic flu prior immunity Substantial impact of missed flu seasons due to the COVID-19 pandemic on prior immunity.* - 50% lower immunity than a typical, pre-COVID19 pandemic season. High Vaccine Effectiveness - VE = 50% against medically attended influenza illnesses and hospitalizations (comparable to 2015-16 season).
Low Vaccine Effectiveness	Scenario C Optimistic flu prior immunity - No impact of missed flu seasons due to the COVID-19 pandemic on prior immunity.* - Same amount of prior immunity as in a typical, pre-COVID19 pandemic prior season. Low Vaccine Effectiveness - VE = 30% against medically attended influenza illnesses and hospitalizations (comparable to 2018-19 season).	Scenario D Pessimistic flu prior immunity Substantial impact of missed flu seasons due to the COVID-19 pandemic on prior immunity.* - 50% lower immunity than a typical, pre-COVID19 pandemic season. Low Vaccination Protection - VE = 30% against medically attended influenza illnesses and hospitalizations (comparable to 2018-19 season).

Combined – COVID-19 and Influenza

Collaboration of multiple academic teams to provide national and state-by-state level projections for 4 aligned scenarios

- COVID-19 Scenarios – Round 16

- | | |
|----------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------|
| <input checked="" type="checkbox"/> Scenario A
High boosters
Moderate immune escape variant
(A-2022-10-29) | <input checked="" type="checkbox"/> Scenario B
High boosters
High immune escape variant
(B-2022-10-29) |
| <input checked="" type="checkbox"/> Scenario C
Low boosters
Moderate immune escape variant
(C-2022-10-29) | <input checked="" type="checkbox"/> Scenario D
Low boosters
High immune escape variant
(D-2022-10-29) |

- Influenza Scenarios – Round 2

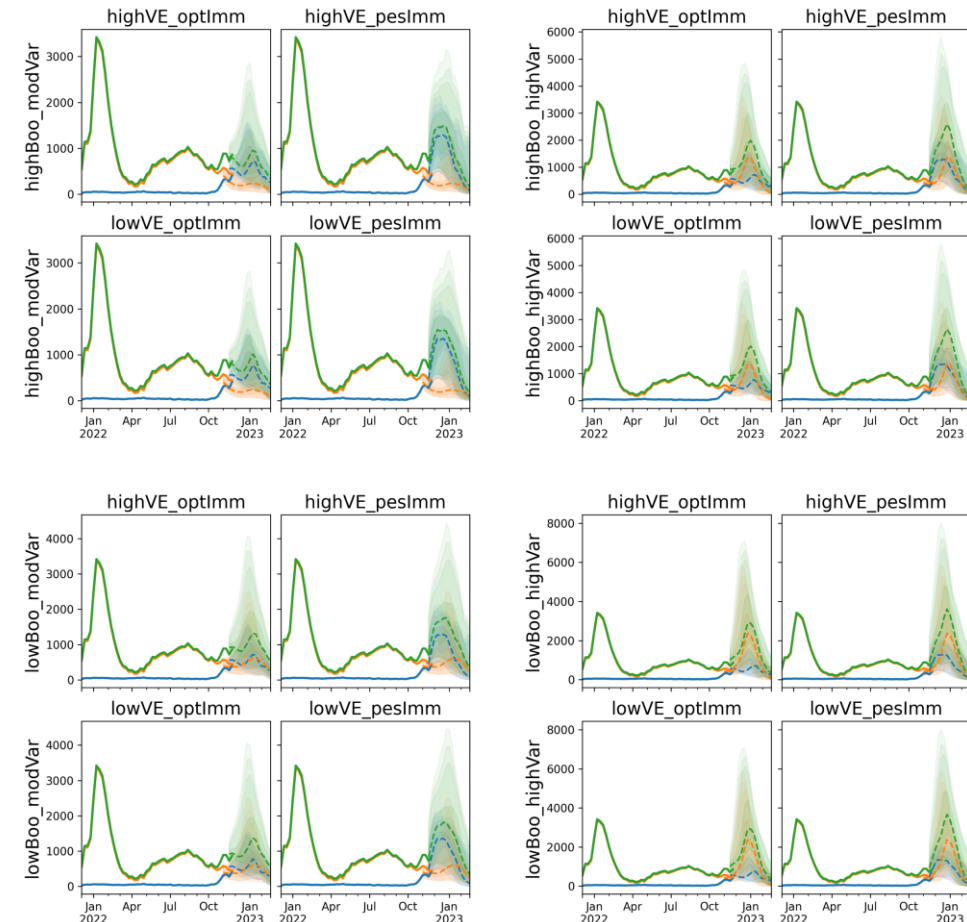
- | | |
|------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------|
| <input checked="" type="checkbox"/> Scenario A
High vaccine effectiveness
optimistic immunity
(A-2022-11-13) | <input checked="" type="checkbox"/> Scenario B
High vaccine effectiveness
pessimistic immunity
(B-2022-11-13) |
| <input checked="" type="checkbox"/> Scenario C
Low vaccine effectiveness
Optimistic immunity
(C-2022-11-13) | <input checked="" type="checkbox"/> Scenario D
Low vaccine effectiveness
pessimistic immunity
(D-2022-11-13) |

Round 16 of COVID-19 is published while Round 2 of Influenza is still in a preliminary result stage

Combined Hospitalizations (VA)

UVA submissions only

VA - Flu Round 2 - COVID Round 16



COVID -19 scenarios and most “pessimistic”
influenza scenarios combined

Key Takeaways

Projecting future cases precisely is impossible and unnecessary.

Even without perfect projections, we can confidently draw conclusions:

- VA weekly case rate is steady at 73/100K from 72/100K
 - US weekly case rate is flat remaining at 76 per 100K from 73 per 100K, though hospitalizations continue a steady rise
- VA hospital occupancy is quickly rising (rolling 7 day mean of 488 from 464 a week ago)
 - Influenza weekly hospital admissions remain high (~300 a week) and are approaching parity with COVID admissions (~40% of the combined)
- Projections anticipate increases in cases and hospitalizations in coming weeks
 - Combined hospitalizations due to Influenza and COVID-19 though are expected to level off
- Model updates:
 - Variant X candidates have now surpassed 50% (BQ.1.1, BN.1, BF.7 and others and XBB among others), 50% remains at Nov 12th
 - Modified Booster Scenarios: Current pace (included in all scenarios) with comparisons between keeping the pace (which is slowing) and a more Pessimistic scenario where vaccination halts at current levels

The situation continues to change. Models continue to be updated regularly.

References

Venkatramanan, S., et al. "Optimizing spatial allocation of seasonal influenza vaccine under temporal constraints." *PLoS Computational Biology* 15.9 (2019): e1007111.

Arindam Fadikar, Dave Higdon, Jiangzhuo Chen, Bryan Lewis, Srinivasan Venkatramanan, and Madhav Marathe. Calibrating a stochastic, agent-based model using quantile-based emulation. *SIAM/ASA Journal on Uncertainty Quantification*, 6(4):1685–1706, 2018.

Adiga, Aniruddha, Srinivasan Venkatramanan, Akhil Peddireddy, et al. "Evaluating the impact of international airline suspensions on COVID-19 direct importation risk." *medRxiv* (2020)

NSSAC. PatchSim: Code for simulating the metapopulation SEIR model. <https://github.com/NSSAC/PatchSim>

Virginia Department of Health. COVID-19 in Virginia. <http://www.vdh.virginia.gov/coronavirus/>

Biocomplexity Institute. COVID-19 Surveillance Dashboard. <https://nssac.bii.virginia.edu/covid-19/dashboard/>

Google. COVID-19 community mobility reports. <https://www.google.com/covid19/mobility/>

Biocomplexity page for data and other resources related to COVID-19: <https://covid19.biocomplexity.virginia.edu/>

Questions?

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